

## SEARCH REQUEST FORM

2-157

Requestor's Name: FONDA Serial Number: 08/462147  
Date: 2-12-96 Phone: 308-1620 Art Unit: 1211

## Search Topic:

Please write a detailed statement of search topic. Describe specifically as possible the subject matter to be searched. Define any terms that may have a special meaning. Give examples or relevant citations, authors keywords, etc., if known. For sequences, please attach a copy of the sequence. You may include a copy of the broadest and/or most relevant claim(s).

JAN

Please search the compositions and  
therapeutic methods of attached  
claims 11, 122, 123, 151, 187,  
216, 218 and 261-264.

Inventors: Rudolf Edgar FALK  
Samuel Simon ASCHLAI

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## STAFF USE ONLY

Date completed: <u>2/15/96</u>	Search Site	Vendors
Searcher: <u>[Signature]</u>	<u>      </u> STIC	<u>      </u> IG Suite
Terminal time: <u>40</u>	<u>  ✓  </u> CM-1	<u>  ✓  </u> STN
Elapsed time: <u>      </u>	<u>      </u> Pre-S	<u>      </u> Dialog
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Total time: <u>60</u>	<u>      </u> N.A. Sequence	<u>      </u> Geninfo
Number of Searches: <u>1</u>	<u>      </u> A.A. Sequence	<u>      </u> SDC
Number of Databases: <u>4</u>	<u>      </u> Structure	<u>      </u> DARC/Questel
	<u>  ✓  </u> Bibliographic	<u>      </u> Other

=> fil embase medline

FILE 'EMBASE' ENTERED AT 14:54:54 ON 13 FEB 96

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FILE 'MEDLINE' ENTERED AT 14:54:54 ON 13 FEB 96

=> d his

(FILE 'HOME' ENTERED AT 14:23:30 ON 13 FEB 96)

SET COST OFF

SET AUHELP OFF

FILE 'REGISTRY' ENTERED AT 14:23:40 ON 13 FEB 96

E HYALURONIC ACID/CN

L1 1 S E3

FILE 'EMBASE' ENTERED AT 14:23:52 ON 13 FEB 96

L2 2359 S L1  
L3 4084 S HYALURONIC ACID/CT OR HYALURONIC ACID DERIVATIVE/CT  
L4 4084 S L2 OR L3  
L5 10014 S IMPLANTATION+NT/CT OR IMPLANT/CT  
L6 554346 S C6.440./CT  
L7 111 S L4 AND L6  
L8 2 S L7 AND L5  
L9 17 S L7 AND E4.80./CT  
L10 284 S L4 AND D20./CT  
L11 80 S L4 AND 037.11./CC  
L12 297 S L10 OR L11  
L13 3 S L12 AND L5  
L14 67 S L12 AND E4.80./CT  
L15 54 S 0186/CT AND (L8 OR L9 OR L12)  
L16 70680 S L6 (L) DT/CT  
L17 4 S L15 AND L16  
L18 6 S L8 OR L17  
L19 53 S L12 AND 0186/CT  
L20 1 S L15 NOT L19  
L21 29 S L3/MAJ AND L19  
L22 2 S L19 AND A18.755./CT  
L23 0 S L19 AND G2.325.800/CT  
L24 9 S L19 AND C2.810./CT  
L25 7 S L19 AND 0001/CT  
L26 19 S L18 OR L22 OR L24 OR L25

FILE 'MEDLINE' ENTERED AT 14:37:54 ON 13 FEB 96

L27 3003 S L1  
L28 4613 S HYALURONIC ACID/CT  
L29 3003 S HYALURONIC ACID/CN  
L30 4613 S L27 OR L28 OR L29  
L31 85793 S IMPLANTS, ARTIFICIAL+NT/CT OR DENTAL IMPLANTATION+NT/CT  
L32 175 S L30 AND L31  
L33 83 S C1./CT AND L30  
L34 2 S L32 AND L33  
L35 10 S C21.866./CT AND L32  
L36 13 S L30 AND D20./CT  
L37 25 S L34 OR L35 OR L36  
L38 22 S L30 AND ADMINISTRATION, TOPICAL+NT/CT  
L39 0 S L33 AND L38  
L40 46 S L37 OR L38

FILE 'MEDLINE, EMBASE' ENTERED AT 14:41:45 ON 13 FEB 96

L41 65 DUP REM L40 L26 (0 DUPLICATES REMOVED)  
E FALK R/AU

FILE 'MEDLINE' ENTERED AT 14:43:02 ON 13 FEB 96

E FALK R/AU

L42 214 S E3,E6  
E ASCULAI S/AU  
L43 6 S E3,E4  
L44 219 S L42 OR L43

L45 2 S L44 AND L30  
 L46 183294 S TRANSPLANTATION+NT/CT  
 L47 104 S L30 AND L46  
 L48 3 S L47 AND C1./CT  
 L49 0 S L47 AND L38  
 L50 0 S L47 AND D20./CT  
 L51 51 S L40 OR L45 OR L48

FILE 'EMBASE' ENTERED AT 14:45:27 ON 13 FEB 96

E FALK R/AU  
 L52 173 S E3,E6  
 E ASCULAI S/AU  
 L53 19 S E3,E4  
 L54 19 S (L52 OR L53) AND L4  
 L55 100 S L4 AND TRANSPLANTATION+NT/CT  
 L56 6 S L55 AND L6  
 L57 11 S L55 AND L12  
 L58 18 S L55 AND 0186/CT  
 L59 51 S L26 OR L54 OR L56 OR L57 OR LL58

FILE 'MEDLINE, EMBASE' ENTERED AT 14:48:22 ON 13 FEB 96

L60 98 DUP REM L51 L59 (4 DUPLICATES REMOVED)

FILE 'EMBASE' ENTERED AT 14:48:44 ON 13 FEB 96

L61 29 S L59 NOT AB/FA  
 L62 22 S L59 NOT L61  
 L63 8 S L61 AND L55  
 L64 30 S L62 OR L63

FILE 'MEDLINE' ENTERED AT 14:50:18 ON 13 FEB 96

L65 17 S L51 NOT AB/FA  
 L66 34 S L51 NOT L65  
 L67 3 S L65 AND L46  
 L68 4 S L65 AND L31  
 L69 3 S L66 AND L46  
 L70 8 S L66 AND L31  
 L71 120 S L30 AND C21.866./CT  
 L72 8 S L71 AND C1./CT  
 L73 24 S L34 OR L35 OR L45 OR L48 OR L67 OR L68 OR L69 OR L70 OR

FILE 'MEDLINE, EMBASE' ENTERED AT 14:54:40 ON 13 FEB 96

L74 51 DUP REM L73 L64 (3 DUPLICATES REMOVED)

FILE 'EMBASE, MEDLINE' ENTERED AT 14:54:54 ON 13 FEB 96

=> d 1-51 cbib ab ct

L74 ANSWER 1 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.  
 95176225 EMBASE Rheumatology. Alarcon G.S.; Straaton K.V.. University  
 of Alabama, Birmingham, AL, United States. Journal of the American  
 Medical Association 273/21 (1721-1723) 1995. ISSN: 0098-7484.  
 CODEN: JAMAAP. Pub. Country: United States. Language: English.  
 Summary Language: English.

AB In some populations, the presence and the dose of the 'rheumatoid'  
 epitope have been associated with severe rheumatoid arthritis. -  
 Newer treatments include oral antibiotics, oral type II  
 collagen, and a number of biological products. - Identification of a  
 mutation in the type II procollagen genes is evidence that some  
 clinical osteoarthritis is genetically related.

CT EMTAGS: diagnosis (0140); therapy (0160); heredity (0137);  
 methodology (0130); automation, computers and data processing  
 (0530); mammal (0738); human (0888); oral drug administration  
 (0181); intraarticular drug administration (0175); topical drug  
 administration (0186); priority journal (0007); review (  
 0001)

Medical Descriptors:

\*rheumatoid arthritis: DI, diagnosis  
 \*rheumatoid arthritis: DT, drug therapy  
 \*gene

\*osteoarthritis: DT, drug therapy  
gene mutation  
quality of life  
systemic lupus erythematosus  
allele  
disease severity  
prognosis  
questionnaire  
arthropathy: DI, diagnosis  
absorptiometry  
fibromyalgia  
pain: DT, drug therapy  
exercise  
bone density  
single photon emission computer tomography  
human  
clinical trial  
oral drug administration  
intraarticular drug administration  
topical drug administration  
priority journal  
review  
Drug Descriptors:  
\*collagen type 2  
\*procollagen: EC, endogenous compound  
\*minocycline: CT, clinical trial  
\*minocycline: AD, drug administration  
\*minocycline: DT, drug therapy  
epitope: EC, endogenous compound  
c reactive protein: EC, endogenous compound  
rheumatoid factor: EC, endogenous compound  
HLA DR4 antigen: EC, endogenous compound  
**antibiotic agent: CT, clinical trial**  
**antibiotic agent: AD, drug administration**  
**antibiotic agent: DT, drug therapy**  
neurotransmitter: EC, endogenous compound  
serotonin: EC, endogenous compound  
substance p: EC, endogenous compound  
capsaicin: PR, pharmaceuticals  
**hyaluronic acid: AD, drug administration**  
nonsteroid antiinflammatory agent: DT, drug therapy  
paracetamol: DT, drug therapy

L74 ANSWER 2 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.  
95073330 EMBASE Effect of sodium hyaluronate on diffuse epithelial  
keratitis after penetrating keratoplasty. Yokoi N.; Yamada J.;  
Nishida K.; Kinoshita S.. Department of Ophthalmology, Kyoto  
Prefectural Univ. of Medicine, Kajicho 465, Kamigyo-ku, Kyoto 602,  
Japan. Transplantation Proceedings 27/1 (1412-1413) 1995. ISSN:  
0041-1345. CODEN: TRPPA8. Pub. Country: United States. Language:  
English.

CT EMTAGS: diagnosis (0140); therapy (0160); mammal (0738); human  
(0888); clinical article (0152); human tissue, cells or cell  
components (0111); aged (0019); adult (0018); topical drug  
administration (0186); priority journal (0007); conference paper  
(0061)

Medical Descriptors:  
\*keratitis: CO, complication  
\*keratitis: DI, diagnosis  
\*keratitis: DT, drug therapy  
medical record  
keratopathy: SU, surgery  
cornea opacity: SU, surgery  
**penetrating keratoplasty**  
staining  
eye photography  
disease severity  
scoring system  
human

clinical article  
human tissue  
aged  
adult  
topical drug administration  
priority journal  
conference paper  
Drug Descriptors:  
**\*hyaluronic acid: DT, drug therapy**  
**ofloxacin: DT, drug therapy**  
steroid: DT, drug therapy  
eye drops: DT, drug therapy

L74 ANSWER 3 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.

95340178 EMBASE Iris ischaemia following penetrating keratoplasty for keratoconus (Urrets- Zavalia syndrome). Tuft S.J.; Buckley R.J.. Moorfields Eye Hospital, City Road, London EC1V 2PD, United Kingdom. Cornea 14/6 (618-622) 1995. ISSN: 0277-3740. CODEN: CORNDB. Pub. Country: United States. Language: English. Summary Language: English.

AB A fixed and dilated pupil is an uncommon postoperative complication after penetrating keratoplasty (PK) for keratoconus. Although the clinical features have been well described, the precise aetiology is uncertain. We performed anterior segment fluorescein angiography in the early postoperative period on three patients who developed fixed, dilated pupils after apparently uncomplicated surgery. All of the eyes had severe iris ischaemia. A possible role for a postoperative rise in intraocular pressure in the aetiology of this syndrome is discussed.

CT EMTAGS: therapy (0160); diagnosis (0140); mammal (0738); human (0888); male (0041); female (0042); case report (0151); adolescent (0017); adult (0018); oral drug administration (0181); intravenous drug administration (0182); topical drug administration (0186); priority journal (0007); article (0060)

Medical Descriptors:

\*iris disease: CO, complication  
**\*penetrating keratoplasty**  
**\*keratoconus: SU, surgery**  
**\*intraocular hypertension: DT, drug therapy**  
**\*ischemia**  
fluorescence angiography  
mydriasis  
glaucoma: DT, drug therapy  
human  
male  
female  
case report  
adolescent  
adult  
oral drug administration  
intravenous drug administration  
topical drug administration  
priority journal  
article  
Drug Descriptors:  
**\*acetazolamide: DT, drug therapy**  
**\*dexamethasone: DT, drug therapy**  
**\*chloramphenicol**  
**\*mydriatic agent: DT, drug therapy**  
**\*hyaluronic acid**  
**\*hydroxypropylmethylcellulose**  
cyclopentolate: DT, drug therapy  
phenylephrine: DT, drug therapy  
mannitol: DT, drug therapy  
procaine: CB, drug combination  
atropine: CB, drug combination  
adrenalin: CB, drug combination  
mydracaine  
unclassified drug

L74 ANSWER 4 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.

95178268 EMBASE Soft tissue augmentation: A review. Elson M.L..  
Dermatology Center, Inc., 4535 Harding Road, Nashville, TN  
37205-2120, United States. Dermatologic Surgery 21/6 (491-502)  
1995. ISSN: 1076-0512. CODEN: DESUFE. Pub. Country: United States.  
Language: English. Summary Language: English.

AB BACKGROUND. Soft tissue augmentation is one of the most sought after  
cosmetic procedures around the world. It has been performed for  
centuries, but only in this decade or so have materials become  
available that allow effective therapy. OBJECTIVE. This review  
article discusses the use of materials for soft tissue augmentation  
as to where they fit into the overall scheme of treating the aging  
face, as well as the benefits and side effects of each, and how to  
maximize the use of these various materials. RESULTS. The materials  
reviewed with regard to both efficacy and safety of those currently  
on the market as well as those currently in research. CONCLUSION.  
With the understanding of the role of soft tissue augmentation in  
the overall treatment of the aging face as well as risk factors and  
ways to minimize the side effects, dermatologic surgeons can reach  
the goal of effectively treating patients with these materials.

CT EMTAGS: diagnosis (0140); apparatus, equipment and supplies (0510);  
therapy (0160); **infection** (0310); mammal (0738); human  
(0888); priority journal (0007); review (0001)

Medical Descriptors:

\*face surgery

\*skin surgery

\*plastic surgery

aging

esthetic surgery

risk factor

scar: SU, surgery

skin test

rhytidectomy

surgical technique

needle

allergic reaction: CO, complication

allergic reaction: DT, drug therapy

cyst: CO, complication

cyst: DT, drug therapy

**abscess: CO, complication**

**abscess: DT, drug therapy**

human

priority journal

review

Drug Descriptors:

\*biomaterial

\*silicone

\*collagen implant

\*atelocollagen

\*politef

\*hyaluronic acid

steroid: DT, drug therapy

antihistaminic agent: DT, drug therapy

antiinflammatory agent: DT, drug therapy

glyceryl trinitrate: DT, drug therapy

dimethyl sulfoxide: DT, drug therapy

L74 ANSWER 5 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.

95336639 EMBASE Lower extremity manifestations of Graves disease. Sauer  
P.; Brandes B.; Mahmarian R.R.. 2001 N. Adams Street, Arlington, VA  
22201, United States. Journal of Foot and Ankle Surgery 34/5  
(489-497) 1995. ISSN: 1067-2516. CODEN: JFSUEI. Pub. Country:  
United States. Language: English. Summary Language: English.

AB Pretibial myxedema (not to be confused with myxedema suggestive of  
hypothyroidism) is one of the extrathyroidal manifestations observed  
in some patients with Graves' disease. Graves' disease is commonly  
described as a disease that consists of one or more of the following  
characteristics: goiter, exophthalmos, acropachy, and pretibial

myxedema. In order to facilitate a better understanding of the pathology and etiology of pretibial myxedema, a review of current research focusing on the immunological basis of Graves' disease is presented. An examination of the signs, symptoms, diagnosis, treatment, and differential diagnosis is also presented, as well as a case study that demonstrates Graves' disease and its extrathyroidal manifestations and complications.

CT EMTAGS: diagnosis (0140); therapy (0160); etiology (0135); mammal (0738); human (0888); nonhuman (0777); female (0042); case report (0151); adult (0018); oral drug administration (0181); topical drug administration (0186); article (0060); adverse drug reaction (0198); iatrogenic disease (0300)

Medical Descriptors:

\*graves disease: DI, diagnosis  
\*graves disease: DT, drug therapy  
\*graves disease: ET, etiology  
\*myxedema: DT, drug therapy  
\*leg ulcer: DI, diagnosis  
\*leg ulcer: DT, drug therapy  
\*leg ulcer: TH, therapy  
\*osteoporosis: DI, diagnosis

biopsy

thyrotoxicosis: DT, drug therapy

x ray

hyperthyroidism: DI, diagnosis

hyperthyroidism: DT, drug therapy

hypothyroidism: SI, side effect

thyroidectomy

radiography

differential diagnosis

bone scintiscanning

debridement

human

nonhuman

female

case report

adult

oral drug administration

topical drug administration

article

Drug Descriptors:

**ciprofloxacin: DT, drug therapy**

**penicillin v: DT, drug therapy**

naproxen: DO, drug dose

naproxen: DT, drug therapy

local anesthetic agent: DT, drug therapy

glycosaminoglycan: EC, endogenous compound

**hyaluronic acid: EC, endogenous compound**

long acting thyroid stimulator: EC, endogenous compound

thyroid hormone: EC, endogenous compound

glucocorticoid: EC, endogenous compound

collagen: EC, endogenous compound

dna: EC, endogenous compound

major histocompatibility antigen: EC, endogenous compound

HLA antigen: EC, endogenous compound

**gamma interferon**

calcium: EC, endogenous compound

phosphate: EC, endogenous compound

**iodine: DT, drug therapy**

**iodine: PD, pharmacology**

propylthiouracil: DT, drug therapy

propylthiouracil: PD, pharmacology

thiamazole: DT, drug therapy

thiamazole: PD, pharmacology

sodium iodide i 131: AE, adverse drug reaction

sodium iodide i 131: DT, drug therapy

steroid: AD, drug administration

steroid: DT, drug therapy

beta adrenergic receptor blocking agent: DT, drug therapy

**polysporin**

mucin: EC, endogenous compound

acetic acid

mannan

## L74 ANSWER 6 OF 51 MEDLINE

95404472 Extracapsular cataract surgery using capsulorhexis with viscoexpression via a limbal section. Burton R L; Pickering S. (Eye Department, West Norwich Hospital, Norfolk, United Kingdom.. ) JOURNAL OF CATARACT AND REFRACTIVE SURGERY, (1995 May) 21 (3) 297-301. Journal code: JPB. ISSN: 0886-3350. Pub. country: United States. Language: English.

AB Two hundred consecutive patients had extracapsular cataract surgery by capsulorhexis and viscoexpression. Capsulorhexis, attempted in 195 eyes, was successful in 87.7%. Viscoexpression was attempted in 162 cases and successfully delivered the nucleus in 87.7%. There were five cases of zonule rupture, one of posterior capsule rupture, and two of vitreous loss. If the capsulorhexis is larger than 5 mm, viscoexpression can be safely used on all cataracts, regardless of nuclear density, and is the ideal transition to phacoemulsification.

CT Check Tags: Human

\*Cataract Extraction: MT, methods

**\*Hyaluronic Acid: AD, administration & dosage**

Intraoperative Complications

\*Lens Capsule, Crystalline: SU, surgery

Lens Nucleus, Crystalline: SU, surgery

**Lenses, Intraocular**

Ligaments: IN, injuries

\*Limbus Corneae: SU, surgery

**Rupture**

Visual Acuity

## L74 ANSWER 7 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.

95357529 EMBASE Liposome-mediated drug targeting in topical and regional therapies. Margalit R.. Department of Biochemistry, Tel Aviv University, Tel Aviv 69978, Israel. Critical Reviews in Therapeutic Drug Carrier Systems 12/2-3 (233-261) 1995. ISSN: 0743-4863. CODEN: CRTSEO. Pub. Country: United States. Language: English. Summary Language: English.

AB Liposome-mediated drug targeting is reviewed in four major categories of topical and regional therapies: wounds and burns, ocular, intraperitoneal, and pulmonary. A survey of the data in the field is preceded by definitions of carrier-mediated drug targeting, in particular for topical and regional treatments. The ability of liposomes to meet essential requirements for task performance and liposome surface-modification as the major approach to endow liposomes with targeting abilities are reviewed. Analysis of current findings in the field shows that (1) most studies explored regular liposomes that were unable to meet the essential requirements for targeting and (2) in vivo drug targeting in topical and regional therapies has been achieved rarely and seldom attempted, yet there are encouraging indications from a few studies that using surface modified liposomes such targeting is feasible. Both established and novel liposomal systems attest to this feasibility and point out future directions. The former can be found by revisiting immunoliposomes that were initially designed for systemic administration but might well fit topical and regional cases. The latter is exemplified by bioadhesive liposomes, designed specifically for topical/regional therapies. It is concluded that careful implementation of such approaches could be successful for the achievement of liposome-mediated drug targeting in topical and regional therapies.

CT EMTAGS: therapy (0160); infection (0310); prevention (0165); injury (0301); skin, hair, nails and sweat glands (0980); topical drug administration (0186); review (0001)

Medical Descriptors:

\*drug targeting

**infection: DT, drug therapy**

infection: PC, prevention



wound: DT, drug therapy  
 burn: DT, drug therapy  
 extracellular matrix  
**skin**  
**sepsis: DT, drug therapy**  
 topical drug administration  
 review  
 Drug Descriptors:  
 \*liposome  
 \*immunoliposome  
 growth factor: AD, drug administration  
 growth factor: DT, drug therapy  
**antibiotic agent: AD, drug administration**  
**antibiotic agent: DT, drug therapy**  
 aerosol  
 drug delivery system  
 neurotransmitter  
 antibody  
 photosensitizing agent  
 photofrin  
 porphyrin: EC, endogenous compound  
 low density lipoprotein: EC, endogenous compound  
 collagen  
 gelatin  
**hyaluronic acid**  
**tobramycin**  
**sulfadiazine silver**  
**cefoxitin: AD, drug administration**  
**cefoxitin: DT, drug therapy**  
**cefazolin**  
**ampicillin**  
**fluconazole**  
 cyclosporin: AD, drug administration  
 cyclosporin: DO, drug dose  
 cyclosporin: PR, pharmaceuticals  
 dna

L74 ANSWER 8 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.

95251913 EMBASE Topical semi-solid drug delivery: Kinetics and tolerance of ophthalmic hydrogels. Zignani M.; Tabatabay C.; Gurny R.. School Pharmacy, University of Geneva, 30 Quai E. Ansermet, CH-1211 Geneve 4, Switzerland. Advanced Drug Delivery Reviews 16/1 (51-60) 1995. ISSN: 0169-409X. CODEN: ADDREP. Pub. Country: Netherlands. Language: English. Summary Language: English.

AB The efficacy of ophthalmic semi-solid hydrogels is mostly based on an increase of ocular residence time, via enhanced viscosity and mucoadhesive properties. Preformed and in particular in situ gelling systems improve bioavailability and decrease the side effects induced by the systemic absorption of topically applied ophthalmic drugs. Since increased viscosity often causes the discomfort of blurred vision and foreign body sensation, it is important to assess the optimal range of viscosity as well as the most suitable rheological behavior which will ensure good efficacy and tolerance.

CT EMTAGS: visual system (0915); pharmacokinetics (0194); mammal (0738); human (0888); nonhuman (0777); topical drug administration (0186); priority journal (0007); review (0001); adverse drug reaction (0198); iatrogenic disease (0300)  
 Medical Descriptors:

\*eye  
 drug administration  
 viscosity  
 drug efficacy  
 drug tolerance  
 drug bioavailability  
 blood rheology  
 eye irritation: SI, side effect  
 visual impairment: SI, side effect  
 human  
 nonhuman

topical drug administration  
priority journal  
review

Drug Descriptors:

\*drug delivery system: PR, pharmaceuticals  
\*hydrogel: AE, adverse drug reaction  
\*hydrogel: PR, pharmaceuticals  
drug vehicle: PR, pharmaceuticals  
pilocarpine: PR, pharmaceuticals  
hydroxyethylcellulose: PR, pharmaceuticals  
polyvinyl alcohol: PR, pharmaceuticals  
**hyaluronic acid: PR, pharmaceuticals**  
carbomer: AE, adverse drug reaction  
carbomer: PR, pharmaceuticals  
cellulose: PR, pharmaceuticals  
**fusidic acid: PK, pharmacokinetics**  
**fusidic acid: PR, pharmaceuticals**  
betaxolol: AE, adverse drug reaction  
betaxolol: PK, pharmacokinetics  
betaxolol: PR, pharmaceuticals  
artificial tear: PR, pharmaceuticals  
unclassified drug  
xanthan: PR, pharmaceuticals  
gellan gum: PR, pharmaceuticals  
poloxamer: AE, adverse drug reaction  
poloxamer: PR, pharmaceuticals  
lacril: PR, pharmaceuticals  
liquifilm tears  
neotears  
polymacon  
hy drop

L74 ANSWER 9 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.

94346252 EMBASE Serum Vpr regulates productive infection and latency of human immunodeficiency virus type 1. Levy D.N.; Refaeli Y.; MacGregor R.R.; Weiner D.B.. Pathology/Laboratory Medicine Dept., University of Pennsylvania, Philadelphia, PA 19104, United States. PROC. NATL. ACAD. SCI. U. S. A. 91/23 (10873-10877) 1994. ISSN: 0027-8424. CODEN: PNASA6. Pub. Country: United States. Language: English. Summary Language: English.

AB In human immunodeficiency virus (HIV)-positive individuals, the vast majority of infected peripheral blood cells and lymph node cells may be latently or nonproductively infected. The vpr open reading frame of HIV-1 encodes a 15-kDa virion-associated protein, Vpr. The vpr gene has been shown to increase virus replication in T cells and monocyte/macrophages in vitro. We have previously reported that vpr expression in various tumor lines leads to growth inhibition and differentiation, indicating that Vpr may function as a regulator of cellular permissiveness to HIV replication. Here we show that Vpr protein is present in significant amounts in the serum of AIDS patients. Purified serum Vpr activated virus expression from five latently infected cell lines, U1, OM.10.1, ACH-2, J1.1, and

LL58. Serum Vpr also activated virus expression from resting peripheral blood mononuclear cells of HIV- infected individuals. Together, these findings implicate serum Vpr in the activation of HIV replication in vivo and in the control of latency. Anti- Vpr antibodies inhibited Vpr activity, suggesting that humoral immunity modulates Vpr activity in vivo. These results have broad implications for the virus life cycle and for the prospective control of HIV replication and pathogenesis.

CT EMTAGS: infection (0310); heredity (0137); blood and hemopoietic system (0927); lymphatic system (0929); virus (0761); mammal (0738); human (0888); human tissue, cells or cell components (0111); priority journal (0007); article (0060)  
Medical Descriptors:  
\*latent virus infection  
\*human immunodeficiency virus infection  
virus gene  
open reading frame

virus replication  
 t lymphocyte  
 human immunodeficiency virus 1  
 gene expression regulation  
 humoral immunity  
 immunoregulation  
 human  
 human cell  
 priority journal  
 article  
 Drug Descriptors:  
 \*vpr protein

L74 ANSWER 10 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.

94351931 EMBASE Local necrosis and fatal perforation of oesophagus after endoscopic ligation [9]. Schoonbroodt D.; Zipf A.; Jung M.. Division of Gastroenterology, Goethe University Hospital, 60590 Frankfurt am Main, Germany, Federal Republic of. LANCET 344/8933 (1365) 1994. ISSN: 0140-6736. CODEN: LANCAO. Pub. Country: United Kingdom. Language: English.

CT EMTAGS: injury (0301); diagnosis (0140); therapy (0160); prevention (0165); **infection** (0310); mammal (0738); human (0888); male (0041); female (0042); case report (0151); adult (0018); priority journal (0007); letter (0008)

Medical Descriptors:

\*esophagus perforation: CO, complication

\*tissue necrosis: CO, complication

\*digestive tract endoscopy

\*ligation

esophagus varices bleeding: SU, surgery

endoscopic sclerotherapy

liver cirrhosis

adrenal tumor

graft rejection: DT, drug therapy

graft rejection: PC, prevention

**kidney transplantation**

esophagus ulcer: CO, complication

esophagus ulcer: DI, diagnosis

**sepsis: DT, drug therapy**

postoperative complication

esophagoscopy

collagen

human

male

female

case report

adult

priority journal

letter

Drug Descriptors:

immunosuppressive agent: DT, drug therapy

immunosuppressive agent: PD, pharmacology

methylprednisolone: CB, drug combination

methylprednisolone: DT, drug therapy

methylprednisolone: PD, pharmacology

cyclosporin: CB, drug combination

cyclosporin: DT, drug therapy

**ceftriaxone: DT, drug therapy**

corticosteroid: DT, drug therapy

corticosteroid: PD, pharmacology

**hyaluronic acid: EC, endogenous compound**

**antibiotic agent**

L74 ANSWER 11 OF 51 MEDLINE

DUPLICATE 1

95012152 Serum hyaluronate in the assessment of liver endothelial cell function after orthotopic liver transplantation in the rat. Shimizu H; He W; Guo P; Dziadkowiec I; Miyazaki M; **Falk R E**.

(Department of Surgery, University of Toronto, Ontario, Canada..

)HEPATOLOGY, (1994 Nov) 20 (5) 1323-9. Journal code: GBZ. ISSN:

0270-9139. Pub. country: United States. Language: English.

AB This study was designed to evaluate the use of serum hyaluronate as a marker of liver endothelial cell function after liver transplantation. We performed orthotopic liver transplantation in both isogeneic and allogeneic rejector models. After transplantation, hepatocyte function was assessed on the basis of serum ALT and total bilirubin levels, and liver endothelial cell function was judged on the basis of serum hyaluronate levels. Significant increase of hyaluronate in the rejector model, compared with the isogeneic model, was seen before any significant results could be obtained from conventional liver function tests. The impaired metabolism of hyaluronate in the rejector model was observed after intravenous injection of trace amounts of radioactive material. Serial studies demonstrate that the endothelial cell is a more susceptible target for the immune response than the hepatocyte. Serum hyaluronate concentration may be a better indicator in the early assessment of graft function. We also examined serum hyaluronate levels to evaluate cold ischemia-reperfusion injury to the liver endothelial cells in the isogeneic model. At 2 hr after reperfusion, hyaluronate levels in the 6-hr cold ischemia (nonviable allograft) group were significantly higher than in the 1-hr and 3-hr cold ischemia (viable allograft) groups. However, there was little difference between the viable allograft groups. After an intravenous injection of 1 mg/kg hyaluronate, the hyaluronate elimination rate in the 3-hr group was distinctly slower than that in the 1-hr group. These data indicate that the hyaluronate elimination rate may be a more sensitive marker of liver endothelial cell function in viable liver after a short period of ischemia.

CT Check Tags: Animal; Male; Support, Non-U.S. Gov't

Cold

Endothelium: PA, pathology

Endothelium: PP, physiopathology

Graft Survival

**\*Hyaluronic Acid: BL, blood**

Ischemia: BL, blood

Ischemia: PA, pathology

Liver: PA, pathology

**\*Liver: PP, physiopathology**

Liver Circulation

**\*Liver Transplantation**

Postoperative Period

Rats

Rats, Inbred ACI

Rats, Inbred Lew

Reperfusion

L74 ANSWER 12 OF 51 MEDLINE

DUPLICATE 2

95169719 Monitoring of acute lung rejection and infection by bronchoalveolar lavage and plasma levels of hyaluronic acid in clinical lung transplantation. Rao P N; Zeevi A; Snyder J; Spichty K; Habrat T; Warty V; Dauber J; Paradis I; Duncan S; Pham S; et al. (Department of Surgery and Pathology, University of Pittsburgh, Pa..) JOURNAL OF HEART AND LUNG TRANSPLANTATION, (1994 Nov-Dec) 13 (6) 958-62. Journal code: A0Q. ISSN: 1053-2498. Pub. country: United States. Language: English.

AB Local immunological injury caused by acute lung rejection leads to fibroblast proliferation. Hyaluronate is a product of activated fibroblasts and possibly an indicator of fibroblast proliferation. One hundred thirty-six bronchoalveolar lavage and plasma hyaluronate assays were performed in 57 lung transplant recipients. Pulmonary endothelial cell function was assessed by measuring bronchoalveolar lavage levels of purine nucleoside phosphorylase. Presence of acute cellular rejection was monitored by transbronchial biopsy histologic evaluation and was classified as minimal to mild (acute rejection I, II) and moderate to severe (acute rejection III, IV). Infection was confirmed by bronchoalveolar lavage culture and antibiotic sensitivity. Bronchoalveolar lavage hyaluronate levels in clinically stable recipients were 33.5 +/- 4.69 micrograms/L and were significantly higher than with clinically stable recipients (p =

0.0001), infection ( $p = 0.008$ ), or mild rejection ( $p = 0.001$ ). Levels were highest in recipients with diffuse alveolar damage ( $392.4 \pm 60.6$  micrograms/L). Diffuse alveolar damage also resulted in significant elevations of plasma HA as compared with stable recipients ( $p = 0.001$ ) and mild rejection. We conclude that clinically significant injury to the allograft from rejection or diffuse alveolar damage can be assessed by bronchoalveolar lavage hyaluronate assays and suggest that the source of hyaluronate in these instances are activated fibroblasts.

CT Check Tags: Female; Human; Male  
 Acute Disease  
 \*Bronchoalveolar Lavage Fluid: CH, chemistry  
 \*Graft Rejection: DI, diagnosis  
 \*Hyaluronic Acid: AN, analysis  
 Hyaluronic Acid: BL, blood  
 \*Infection: DI, diagnosis  
 Infection: ET, etiology  
 Lung Diseases: DI, diagnosis  
 \*Lung Transplantation  
 \*Postoperative Complications: DI, diagnosis  
 Purine-Nucleoside Phosphorylase: AN, analysis

L74 ANSWER 13 OF 51 MEDLINE DUPLICATE 3  
 94378312 Evaluation of preservation damage to liver endothelial cells by hyaluronic acid uptake in vitro. Shimizu H; He W; Guo P; Miyazaki M; Falk R E. (Department of Surgery, University of Toronto, Ontario, Canada.. )TRANSPLANTATION, (1994 Sep 15) 58 (5) 635-6. Journal code: WEJ. ISSN: 0041-1337. Pub. country: United States. Language: English.

CT Check Tags: Animal; In Vitro; Male  
 \*Cryopreservation  
 Endothelium: CY, cytology  
 Endothelium: ME, metabolism  
 Evaluation Studies  
 \*Hyaluronic Acid: PK, pharmacokinetics  
 \*Liver: ME, metabolism  
 Liver Transplantation  
 \*Organ Preservation  
 Rats  
 Rats, Inbred Lew  
 Time Factors

L74 ANSWER 14 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.  
 94268137 EMBASE Suppression of corneal allograft rejection by systemic cyclosporine-A in heavily vascularized rabbit corneas following alkali burns. Rehany U.; Waisman M.. Department of Ophthalmology, Western Galilee Medical Center, P.O.B. 21, Nahariya 22100, Israel. CORNEA 13/5 (447-453) 1994. ISSN: 0277-3740. CODEN: CORNDB. Pub. Country: United States. Language: English. Summary Language: English.

AB Immunologic rejection is the main cause of corneal graft failure, especially in vascularized corneal beds. The purpose of this study was to investigate the effect of systemic Cyclosporine-A (CsA) on the survival of corneal allografts in heavily vascularized rabbit corneal beds, following alkali burn. Heavy corneal vascularization was induced in one eye of 20 rabbits by alkali burn. Forty-five days later, penetrating keratoplasty was performed in all the heavily vascularized corneas. Twenty-five mg/kg/day of CsA was intramuscularly administered to 10 rabbits for 30 days. The other 10 rabbits were treated with the solvent without CsA and were used as a matched control group. The results show a significant difference in corneal allograft survival between the two groups. All corneal grafts in the untreated group were intensely rejected and vascularized within 3 weeks. Nine of the 10 corneal transplants, in the CsA-treated group, remained transparent without signs of immunologic rejection for >180 days. In one corneal transplant, minor signs of rejection occurred. We suggest that CsA, when given systemically, is a potent drug in the prevention of immunologic rejection in high-risk corneal transplantations, such as allografts,

in heavily vascularized corneas following alkali burn.  
 CT EMTAGS: diagnosis (0140); therapy (0160); prevention (0165); injury (0301); rabbits and hares (0731); mammal (0738); nonhuman (0777); animal experiment (0112); animal model (0106); biological model (0502); controlled study (0197); animal tissue, cells or cell components (0105); intramuscular drug administration (0184); intravenous drug administration (0182); topical drug administration (0186); priority journal (0007); article (0060)

Medical Descriptors:

**\*cornea graft**

\*graft rejection: CO, complication  
 \*graft rejection: DI, diagnosis  
 \*graft rejection: DT, drug therapy  
 \*graft rejection: PC, prevention  
 \*cornea neovascularization: CO, complication  
 \*cornea neovascularization: SU, surgery  
 \*cornea burn  
 \*caustic burn  
 immunosuppressive treatment

**graft survival**

**allograft**

**penetrating keratoplasty**

**rabbit**

**cornea transplantation**

**high risk patient**

**nonhuman**

**animal experiment**

**animal model**

**controlled study**

**animal tissue**

**intramuscular drug administration**

**intravenous drug administration**

**topical drug administration**

**priority journal**

**article**

Drug Descriptors:

**\*cyclosporin a: DT, drug therapy**

**sodium hydroxide**

**pentobarbital**

**oxybuprocaine**

**cyclopentolate**

**chloramphenicol**

**ointment**

**benzathine penicillin**

**hyaluronic acid**

**heparin**

**castor oil**

**atropine**

L74 ANSWER 15 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.

94379806 EMBASE Postoperative management of corneal graft. Saini J.S..  
 Postgraduate Inst. Med. Educat./Res., Chandigarh 160 012, India.  
 Indian Journal of Ophthalmology 42/4 (215-217) 1994. ISSN:  
 0301-4738. CODEN: IJOMBM. Pub. Country: India. Language: English.

CT EMTAGS: therapy (0160); prevention (0165); mammal (0738); human (0888); article (0060)

Medical Descriptors:

**\*cornea transplantation**

**follow up**

**postoperative care**

**intraocular pressure**

**graft rejection: PC, prevention**

**postoperative complication**

**human**

**article**

Drug Descriptors:

**antibiotic agent**

**cyclopentolate**

**acetazolamide**

**hyaluronic acid**  
corticosteroid

- L74 ANSWER 16 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.  
95308113 EMBASE Effects of hyaluronic acid on experimental tumor uptake of 5-fluorouracil. Klein E.S.; He W.; Shmizu S.; **Asculai S.**; **Falk R.E.**; Ben-Ari G.Y.. Department of Surgery C, Chaim Sheba Medical Center, Tel-Hashomer 52621, Israel. Regional Cancer Treatment 7/3-4 (163-164) 1994. ISSN: 0935-0411. CODEN: RCTRED. Pub. Country: Germany, Federal Republic of. Language: English. Summary Language: English.
- AB Biological and chemical properties of hyaluronic acid (HA) qualify this macromolecule as a prospective carrier of drugs to various organs. We studied the effects of HA on tritiated 5-Fluorouracil uptake (3H-5-FU) by various experimental tumor models. Three groups of rats were studied: A - with liver implanted rat mammary Ca (RMC), B and C - with subcutaneous Fisher bladder Ca (FBC). Groups A and B received IV 3H-5-FU alone or combined with HA. Group C received intratumoral 3H-5-FU either with or without HA. Uptake and retention of 3H-5-FU in the tumors and in normal liver and skin tissues (controls) was measured at various time intervals. Uptake of 3H-5-FU combined with HA by tumor tissue was significantly higher ( $p < 0.05$ ) than that of 5-FU alone. Retention of 5-FU combined with HA in tumor tissue is higher than in non-tumorous controls: after 6 hours, 85% of 5-FU is retained in the tumors, whereas only 50% in the controls ( $p > 0.05$  by ANOVA). These results suggest that HA may effect 3H-5-FU uptake and retention in the rat tumor model.
- CT EMTAGS: malignant neoplastic disease (0306); digestive system (0935); liver (0946); skin, hair, nails and sweat glands (0980); nonhuman (0777); rat (0733); mammal (0738); controlled study (0197); animal experiment (0112); animal model (0106); biological model (0502); intravenous drug administration (0182); intradermal drug administration (0176); priority journal (0007); article (0060); therapy (0160); pharmacokinetics (0194); radioisotope (0131)
- Medical Descriptors:  
\*liver tumor  
\*subcutaneous tissue tumor  
drug uptake  
cancer graft  
breast carcinoma  
liver  
bladder carcinoma  
skin  
nonhuman  
rat  
controlled study  
animal experiment  
animal model  
intravenous drug administration  
intradermal drug administration  
priority journal  
article
- Drug Descriptors:  
\*fluorouracil: IT, drug interaction  
\*fluorouracil: CB, drug combination  
\*fluorouracil: PK, pharmacokinetics  
\*hyaluronic acid: IT, drug interaction  
\*hyaluronic acid: CB, drug combination  
drug carrier  
radioisotope

- L74 ANSWER 17 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.  
94280194 EMBASE [The medical-surgical treatment of the bet sore (II)]. TRATAMIENTO MEDICO-QUIRURGICO DE LAS ULCERAS POR DECUBITO (II). Martin Bertolin S.; Gonzalez Martinez R.; Garay Burdeos M.; Neira Gimenez C.; Marquina Vila P.; Amorrortu Velayos J.. Unid. de Cirugia Plastica/Reparadora, Hospital General Universitario, Avda. Tres Cruces, s/n, 46014 Valencia, Spain. CIENC. PHARM. 4/3 (137-143) 1994. ISSN: 1131-5253. CODEN: CIPHEA. Pub. Country: Spain. Language:

Spanish. Summary Language: English; Spanish.

AB Pressure sores are particularly common in the elderly. Both medical and surgical treatments are possible, depending on a number of factors. The authors present their experience with different approaches.

CT EMTAGS: injury (0301); therapy (0160); apparatus, equipment and supplies (0510); mammal (0738); human (0888); topical drug administration (0186); review (0001); enzyme (0990)

Medical Descriptors:

**\*decubitus: TH, therapy**

**\*decubitus: DT, drug therapy**

**\*ulcer: TH, therapy**

**\*ulcer: DT, drug therapy**

drug information

drug mechanism

drug indication

tissue adhesive

drug choice

human

topical drug administration

review

Drug Descriptors:

**\*proteinase: PD, pharmacology**

**\*proteinase: DT, drug therapy**

**\*calcium alginate: PD, pharmacology**

**\*calcium alginate: DT, drug therapy**

**\*sulfadiazine silver: PD, pharmacology**

**\*sulfadiazine silver: DT, drug therapy**

**\*dextranomer: PD, pharmacology**

**\*dextranomer: DT, drug therapy**

**\*cadexomer iodine: PD, pharmacology**

**\*cadexomer iodine: DT, drug therapy**

**antibiotic agent: DT, drug therapy**

**hyaluronic acid: DT, drug therapy**

**solcoseryl: DT, drug therapy**

**oxaceprol: DT, drug therapy**

**acexamic acid: DT, drug therapy**

**silicone: DT, drug therapy**

**zinc oxide: DT, drug therapy**

**pandermin**

**polyurethan**

**nitrofuril: DT, drug therapy**

**silidermil**

**unclassified drug**

**proskin**

L74 ANSWER 18 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.

94223102 EMBASE [Treatment of psoriatic onychodystrophy with a hyaluronic acid product and chondroitinsulphates]. TRATTAMENTO DELL'ONICODISTROFIA PSORIASICA CON UN PRODOTTO A BASE DI ACIDO IALURONICO E CONDROITINSOLFATI. Flori M.L.; Rubegni P.; Micheli S.; Andreassi L.. Istituto Clinica Dermosifilopatica, Università degli Studi, Viale Bracci, 53100 Siena, Italy. G. ITAL. DERMATOL. VENEREOL. 129/3 (129-133) 1994. ISSN: 0026-4741. CODEN: GIDVDZ. Pub. Country: Italy. Language: Italian. Summary Language: Italian; English.

AB The efficacy of a product containing hyaluronic acid and chondroitin sulphates was tested in a double-blind study versus placebo in 30 patients with psoriatic onychodystrophy. The patients were divided into two groups of 15 patients treated with the product and placebo respectively. An improvement with respect to controls was noted in patients using the product. The differences were significant for onychorrexia, onycholysis and subungueal hyperkeratosis. The results may be due to the hydrophilic property of hyaluronic acid and chondroitin sulphates, or perhaps even a direct effect on nail growth.

CT EMTAGS: therapy (0160); mammal (0738); human (0888); controlled study (0197); clinical article (0152); human experiment (0104);



topical drug administration (0186); article (0060)

Medical Descriptors:

**\*psoriasis**

**\*nail dystrophy: DT, drug therapy**

**\*nail dystrophy: CO, complication**

human

controlled study

clinical article

clinical trial

topical drug administration

article

Drug Descriptors:

**\*hyaluronic acid derivative: DT, drug therapy**

**\*hyaluronic acid derivative: CB, drug combination**

**\*hyaluronic acid derivative: CT, clinical trial**

**\*chondroitin sulfate: DT, drug therapy**

**\*chondroitin sulfate: CB, drug combination**

**\*chondroitin sulfate: CT, clinical trial**

**\*retinol: DT, drug therapy**

**\*retinol: CB, drug combination**

**\*retinol: CT, clinical trial**

**\*pyridoxine: DT, drug therapy**

**\*pyridoxine: CB, drug combination**

**\*pyridoxine: CT, clinical trial**

**\*alpha tocopherol: DT, drug therapy**

**\*alpha tocopherol: CB, drug combination**

**\*alpha tocopherol: CT, clinical trial**

placebo

betamethasone: DT, drug therapy

betamethasone: CB, drug combination

**kanamycin: DT, drug therapy**

**kanamycin: CB, drug combination**

unclassified drug

kevis nails: DT, drug therapy

kevis nails: CT, clinical trial

L74 ANSWER 19 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.

94243059 EMBASE Corneal complications of cataract surgery. Green W.T.;

Muir M.G.K.. Department of Ophthalmology, St. Thomas' Hospital,

London SE1 7EH, United Kingdom. CURR. OPIN. OPHTHALMOL. 5/4

(98-104) 1994. ISSN: 1040-8738. CODEN: COOTEF. Pub. Country: United States. Language: English. Summary Language: English.

AB Developments in cataract surgery have stimulated a greater interest

in minimizing unwanted effects of cataract surgery on the cornea.

The two main areas of concern are protection of the corneal endothelium and minimizing distortion of the anterior corneal surface. Endothelial cell loss is of particular importance where there is a preexisting significantly low cell count due to ocular trauma, surgery, or dystrophy, and in situations where cataract extraction is combined with other procedures that may be prolonged or require extensive manipulation. Recent availability of heavier molecular- weight viscoelastic substances are expected to help in minimizing endothelial cell trauma in these situations. In terms of postoperative corneal astigmatism the emphasis has changed for those who are regularly performing phacoemulsification from minimizing surgically induced astigmatism to planning the procedure so that it incorporates a correction of preexisting astigmatism. This emphasis may be more significant in cases of previous anterior segment surgery or trauma.

CT EMTAGS: visual system (0915); apparatus, equipment and supplies (0510); automation, computers and data processing (0530); therapy (0160); **infection** (0310); injury (0301); mammal (0738); human (0888); nonhuman (0777); priority journal (0007); article (0060)

Medical Descriptors:

**\*cornea**

**\*cataract: SU, surgery**

**\*cornea endothelium**

**\*astigmatism: CO, complication**

\*astigmatism: SU, surgery  
 cell density  
 cell loss  
 phacoemulsification  
 specular microscopy  
 lens implant  
 cell structure  
 photometry  
 electric potential  
 computer  
 morphometrics  
 keratopathy: CO, complication  
 cornea edema: CO, complication  
 cornea edema: DT, drug therapy  
**eye infection: CO, complication**  
 cornea perforation: CO, complication  
 cornea perforation: SU, surgery  
**cornea transplantation**  
 fluorophotometry  
 human  
 nonhuman  
 priority journal  
 article  
 Drug Descriptors:  
 free radical: TO, drug toxicity  
**antibiotic agent**  
 acetylcholine  
 adrenalin  
**hyaluronic acid: EC, endogenous compound**  
 chondroitin sulfate  
 hydroxymethylcellulose  
**hydrogen peroxide**  
 ascorbic acid  
 oxygen  
 bicarbonate: EC, endogenous compound  
 dexamethasone: DT, drug therapy  
 prednisolone acetate: DT, drug therapy  
 collagen

L74 ANSWER 20 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.

94243056 EMBASE Corneal storage and complications related to grafting.  
 Ehlers N.; Hjortdal J.; Moller-Pedersen T.. Department of  
 Ophthalmology, Aarhus University Hospital, DK-8000 Aarhus, Denmark.  
 CURR. OPIN. OPHTHALMOL. 5/4 (75-80) 1994. ISSN: 1040-8738. CODEN:  
 COOTEF. Pub. Country: United States. Language: English. Summary  
 Language: English.

AB This review covers the literature during the past year and follows  
 up results published on corneal storage techniques and complications  
 related to corneal grafting. It considers the recent progress and  
 suggests new perspectives on the reconstituted or renovated human  
 donor cornea. It might be possible to revive postmortem donor  
 corneas with new cells, eg, epithelial, endothelial, or keratocytes,  
 drawn from the future recipient or eventually with transgenic  
 multidonor cells. The future holds promise for the development of a  
 new concept in corneal banking, where we leave the period of  
 conservation and enter the era of donor cornea production.

CT EMTAGS: blood and hemopoietic system (0927); cell, tissue or organ  
 culture (0103); visual system (0915); therapy (0160); prevention  
 (0165); immunological procedures (0102); mammal (0738); human  
 (0888); nonhuman (0777); intraperitoneal drug administration (0178);  
 priority journal (0007); review (0001); enzyme (0990)

Medical Descriptors:

**\*cornea graft**  
 cornea preservation  
 cryopreservation  
 freezing  
 thawing  
 serum  
 organ culture

cornea endothelium  
 cornea epithelium  
 cornea cell  
 microscopy  
 graft rejection: CO, complication  
 graft rejection: DT, drug therapy  
 graft rejection: PC, prevention  
 HLA typing  
 immunosuppressive treatment  
**graft survival**  
 cell density  
 cell viability  
 human  
 nonhuman  
 intraperitoneal drug administration  
 priority journal  
 review  
 Drug Descriptors:  
**antibiotic agent**  
 dextran  
 platelet derived growth factor  
 dna: EC, endogenous compound  
 retinol  
 protein kinase c: EC, endogenous compound  
 urokinase: EC, endogenous compound  
**hyaluronic acid: EC, endogenous compound**  
 steroid: AD, drug administration  
 cyclosporin: DT, drug therapy  
 HLA antigen class 1: EC, endogenous compound  
 tsukubaenolide: AD, drug administration  
 tsukubaenolide: DT, drug therapy  
 liposome  
 epidermal growth factor  
 interleukin 1

L74 ANSWER 21 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.

94338422 EMBASE Review and evaluation of 3% diclofenac in hyaluronan (D.HA) gel. Russell A.L.; Fraser R.; Willoughby D.; Tomlinson A.;

**Falk R.E.** Academy of Pain Management, 18 Kensington Road, Bramalea, Ont. L6T 4S5, Canada. ROUND TABLE SER. R. SOC. MED. -/33 (64-71) (1994). ISSN: 0268-3091. CODEN: RTSSES. Pub. Country: United Kingdom. Language: English. Summary Language: English.

AB 1. D.HA has a unique analgesic action distal from the site of inflammation. 2. Consideration should be given in further trials to extending the age group limit to 75 to cover the cases where topical agents will be most useful. 3. Possible evaluation and double blind study for treatment of thrombophlebitis should be undertaken in an older age group who are at higher risk from oral NSAIDs. 4. Capsaicin should be scientifically evaluated as a test bed for rapid inexpensive evaluation of D.HA, and seems to be ideal for comparison with other NSAIDs. Further work is needed in a university laboratory setting. 5. With the ever-increasing epidemic of myofascial and fibromyalgia, thought should be given to evaluating treatment in this field. In summary, HA in combination with an NSAID will induce local analgesia, and distant analgesia in deeper structures beyond the range of initial diffusion. Can this be explained by an axon reflex? Comments would be appreciated.

CT EMTAGS: therapy (0160); nervous system (0910); injury (0301); mammal (0738); human (0888); nonhuman (0777); topical drug administration (0186); human experiment (0104); conference paper (0061)

Medical Descriptors:

\*analgesia  
 drug formulation  
 pain: DT, drug therapy  
 inflammation: DT, drug therapy  
 thrombophlebitis: DT, drug therapy  
 myofascial pain: DT, drug therapy  
 fibromyalgia: DT, drug therapy  
 nerve fiber

soft tissue injury: DT, drug therapy  
 osteoarthritis: DT, drug therapy  
 neuritis: DT, drug therapy  
 ulcer: DT, drug therapy  
 thermography  
 drug mechanism  
 nerve ending  
 nerve stimulation  
 tooth extraction  
 antiinflammatory activity  
 patient compliance  
 human  
 nonhuman  
 topical drug administration  
 clinical trial  
 meta analysis  
 conference paper  
 Drug Descriptors:  
 \*diclofenac: CT, clinical trial  
 \*diclofenac: CM, drug comparison  
 \*diclofenac: DT, drug therapy  
 \*diclofenac: PR, pharmaceuticals  
 \*diclofenac: PD, pharmacology  
**hyaluronic acid: CB, drug combination**  
**hyaluronic acid: DT, drug therapy**  
 nonsteroid antiinflammatory agent: CM, drug comparison  
 capsaicin  
 antibiotic agent: CB, drug combination  
 antibiotic agent: DT, drug therapy  
 substance p: EC, endogenous compound  
 piroxicam: CT, clinical trial  
 piroxicam: CB, drug combination  
 piroxicam: DT, drug therapy

L74 ANSWER 22 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.  
 94338415 EMBASE Effect of hyaluronic acid on the penetration and  
 targeting of drugs. **Falk R.** Department of Surgery, Falk  
 Oncology Centre, 890 Yonge Street, Toronto, Ont. M4W 3PE, Canada.  
 ROUND TABLE SER. R. SOC. MED. -/33 (2-10) 1994 ISSN: 0268-3091.  
 CODEN: RTSSES. Pub. Country: United Kingdom. Language: English.  
 CT EMTAGS: pharmacokinetics (0194); therapy (0160); malignant  
 neoplastic disease (0306); lymphatic system (0929); mammal (0738);  
 human (0888); nonhuman (0777); intravenous drug administration  
 (0182); human experiment (0104); conference paper (0061)  
 Medical Descriptors:  
 \*drug penetration  
 \*drug targeting  
 cancer: DT, drug therapy  
 brain edema: DT, drug therapy  
 extracellular matrix  
 drug binding  
 pathophysiology  
**liver transplantation**  
 drug contraindication  
 lymph vessel  
**graft survival**  
 pain: DT, drug therapy  
 breast cancer: DT, drug therapy  
 breast cancer: RT, radiotherapy  
 artery disease: DT, drug therapy  
 human  
 nonhuman  
 intravenous drug administration  
 clinical trial  
 meta analysis  
 conference paper  
 Drug Descriptors:  
 \*hyaluronic acid  
 nonsteroid antiinflammatory agent: CT, clinical trial

nonsteroid antiinflammatory agent: CB, drug combination  
 nonsteroid antiinflammatory agent: DT, drug therapy  
 nonsteroid antiinflammatory agent: PR, pharmaceuticals  
 diclofenac: CT, clinical trial  
 diclofenac: DT, drug therapy  
 diclofenac: PR, pharmaceuticals  
 cyclosporin: DV, drug development  
 cyclosporin: PR, pharmaceuticals  
 ketorolac: CT, clinical trial  
 ketorolac: CB, drug combination  
 ketorolac: DT, drug therapy  
 ketorolac: PR, pharmaceuticals  
 ascorbic acid: CT, clinical trial  
 ascorbic acid: CB, drug combination  
 ascorbic acid: DT, drug therapy  
 methotrexate: CB, drug combination  
 methotrexate: DT, drug therapy  
 fluorouracil: CB, drug combination  
 fluorouracil: DT, drug therapy  
 tamoxifen: DT, drug therapy  
 mitoxantrone: CB, drug combination  
 mitoxantrone: DT, drug therapy  
 mitomycin c: CB, drug combination  
 mitomycin c: DT, drug therapy  
 edetic acid: CT, clinical trial  
 edetic acid: CB, drug combination  
 edetic acid: DT, drug therapy

L74 ANSWER 23 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.

93141735 EMBASE Functional studies on the human transplanted small  
 intestine. Knutson L.; Meurling S.; Wahlberg J.; Ewald O.; Hallgren  
 R.; Tufveson G.. Dept of Surgery, University Hospital, S-75185  
 Uppsala, Sweden. CLIN. TRANSPLANT. 7/2 (151-157) 1993. ISSN:  
 0902-0063. CODEN: CLTRED. Pub. Country: Denmark. Language: English.  
 Summary Language: English.

AB Intestinal transplantation is currently under development for  
 treatment of short bowel syndrome. It was our purpose to examine the  
 release of inflammatory mediators in a 1-year-old child with  
 congenital aganglionosis following small intestinal transplantation  
 (ligament of Treitz to colon; ileostomy). After obtaining  
 institutional and parental consent, studies were performed at the  
 6th, 7th and 8th week after surgery and while the patient was  
 receiving enteral nutrition (human breast milk). Prior to each  
 investigation the ileal mucosa was examined endoscopically and  
 biopsies were obtained. A 3-cm segment of ileum was isolated between  
 balloons and perfused with an isoosmolar solution at 2 ml/min at  
 37.degree.C. Effluents were analyzed for: albumin, histamine,  
 hyaluronan (hyaluronic acid), eosinophilic cationic protein (ECP)  
 and PGE2. Although control values were unavailable in infants, data  
 obtained from the adult jejunum served as control. Endoscopically  
 the mucosa revealed loss of valvulae conniventes and progressive  
 aphthous to deep mucosal ulcerations. Histologically this was  
 detected as increased number of mononuclear cells, edema and  
 fibrosis. After transplantation the appearance rate of histamine and  
 ECP were low, but increased progressively. PGE2 was markedly  
 increased at the start of the studies (5566pg/cm/h compared to 11.7  
 +- 3.0 in controls, mean +- SEM; n = 35), but did not profoundly  
 increase further. Albumin and particularly hyaluronan, however,  
 increased more than 30-fold versus controls (21420 .mu.g/cm/h vs  
 controls 669 +- 46, n = 66 and 20558ng/cm/h vs controls 660 +-  
 44, n = 66, respectively) in line with clinical deterioration. We  
 conclude: 1) The technique for segmental intestinal perfusion can be  
 used to monitor inflammatory mediators and mucosal leakage in a  
 small bowel allograft. 2) The transplanted small intestine undergoes  
 progressive ulceration associated with histological fibrosis; and,  
 3) The excessive accumulation of interstitial hyaluronan can  
 influence water transport and thereby also intestinal circulation.

CT EMTAGS: congenital disorder (0315); therapy (0160); digestive system  
 (0935); small intestine (0941); etiology (0135); reticuloendothelial

system (0924); **infection** (0310); plant (0699); fungus (0763); mammal (0738); human (0888); female (0042); case report (0151); controlled study (0197); human tissue, cells or cell components (0111); infant (0014); child (0022); priority journal (0007); article (0060)

Medical Descriptors:

**\*intestine transplantation**

short bowel syndrome: SU, surgery  
 intestine secretion  
 digestive system inflammation  
 aganglionosis: CN, congenital disorder  
 ileostomy  
 enteric feeding  
 ileum mucosa  
 endoscopic biopsy  
 ileum  
 tissue perfusion  
 jejunum  
 intestine ulcer: ET, etiology  
 mononuclear cell  
 edema  
 fibrosis  
 drug tissue level  
 mediator

**allotransplantation**

intestine blood flow  
 immunosuppressive treatment  
 graft rejection: CO, complication  
 graft rejection: DT, drug therapy  
 water loss  
 serosa

**septicemia**

candida albicans  
 heart abscess  
 intestine perfusion  
 mucosa cell  
 human  
 female  
 case report  
 controlled study  
 human tissue  
 infant  
 priority journal  
 article

Drug Descriptors:

breast milk  
 albumin: EC, endogenous compound  
 histamine: EC, endogenous compound  
**hyaluronic acid: EC, endogenous compound**  
 protein: EC, endogenous compound  
 prostaglandin e2: EC, endogenous compound  
 prednisone: DT, drug therapy  
 azathioprine: DT, drug therapy  
 thymocyte antibody: DT, drug therapy  
 15 deoxyspergualin: DT, drug therapy  
 okt 3: DT, drug therapy

L74 ANSWER 24 OF 51 MEDLINE

92393965 Hyaluronan: relationship to hemodynamics and survival in porcine injury and sepsis. Berg S; Jansson I; Hesselvik F J; Laurent T G; Lennquist S; Walther S. (Department of Anesthesiology, University Hospital, Linköping, Sweden.. ) CRITICAL CARE MEDICINE, (1992 Sep) 20 (9) 1315-21. Journal code: DTF. ISSN: 0090-3493. Pub. country: United States. Language: English.

AB BACKGROUND AND METHODS: Hyaluronan is a polysaccharide normally present in low concentrations in the blood, and is rapidly cleared from the blood by the liver. Increased plasma hyaluronan concentrations have been found in patients with sepsis. We studied changes in serum hyaluronan concentrations and their relationship to

hemodynamics and survival in a 48-hr porcine model of injury and sepsis. RESULTS: Circulating hyaluronan concentrations increased to high values after induction of experimental sepsis (from mean baseline values of 242 +/- 26 [SEM] to mean maximum concentrations of 964 +/- 255 micrograms/L [p less than .01]) compared with controls (199 +/- 38 to 303 +/- 32 micrograms/L). A weak negative correlation between mean arterial pressure (MAP) and serum hyaluronan values was found ( $r^2 = .47$ ; p less than .01). Nonsurvivors had higher mean serum hyaluronan concentrations than survivors (603 +/- 147 vs. 285 +/- 43 micrograms/L [p less than .05]). CONCLUSIONS: Experimental sepsis is associated with an increase in serum hyaluronan values. The relationship between decreased MAP and increased serum hyaluronan concentrations could point to reduced liver perfusion as a cause. An association between high hyaluronan values and nonsurvival in sepsis is possible.

CT Check Tags: Animal; Comparative Study; Support, Non-U.S. Gov't

Analysis of Variance

Disease Models, Animal

\*Femoral Fractures: BL, blood

Femoral Fractures: MO, mortality

Femoral Fractures: PP, physiopathology

Hemodynamics

\*Hyaluronic Acid: BL, blood

\*Staphylococcal Infections: BL, blood

Staphylococcal Infections: MO, mortality

Staphylococcal Infections: PP, physiopathology

Swine

\*Swine Diseases: BL, blood

Swine Diseases: MO, mortality

Swine Diseases: PP, physiopathology

Time Factors

\*Wounds, Gunshot: BL, blood

Wounds, Gunshot: MO, mortality

Wounds, Gunshot: PP, physiopathology

L74 ANSWER 25 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.

92252432 EMBASE [Wound care]. WUNDVERSORGUNG. Gresser J.; Bitz K.; Heggin J.. Spital Richterswil, Bergstrasse 16, CH-8805 Richterswil, Switzerland. THER. UMSCH. 49/7 (423-428) 1992. ISSN: 0040-5930. CODEN: THUMAM. Pub. Country: Switzerland. Language: German. Summary Language: English; German; French.

AB The following article is a check-list for wound care giving some practical hints. Special interest has been given to the themes of local anesthesia and prevention of infections. The indications and limits of the ambulant wound care are also discussed. Finally, a short explanation is given for the treatment of wounds situated at delicate regions of the body.

CT EMTAGS: injury (0301); therapy (0160); prevention (0165);

infection (0310); classification (0520); mammal (0738);

human (0888); intramuscular drug administration (0184); intravenous drug administration (0182); topical drug administration (

0186); transdermal drug administration (0285); short survey

(0002); adverse drug reaction (0198); iatrogenic disease (0300);

apparatus, equipment and supplies (0510)

Medical Descriptors:

\*wound care

\*wound: DT, drug therapy

\*wound: SU, surgery

drug mixture

prophylaxis

anamnesis

wound infection: PC, prevention

wound infection: DT, drug therapy

disease classification

drug efficacy

toxicity: SI, side effect

allergic reaction: SI, side effect

human

intramuscular drug administration

intravenous drug administration  
 topical drug administration  
 transdermal drug administration  
 short survey

Drug Descriptors:

**\*antiinfective agent: DT, drug therapy**

procaine: AE, adverse drug reaction  
 procaine: CM, drug comparison  
 procaine: DT, drug therapy  
 lidocaine: AE, adverse drug reaction  
 lidocaine: CM, drug comparison  
 lidocaine: DT, drug therapy  
 mepivacaine: AE, adverse drug reaction  
 mepivacaine: CM, drug comparison  
 mepivacaine: DT, drug therapy  
 bupivacaine: AE, adverse drug reaction  
 bupivacaine: CM, drug comparison  
 bupivacaine: DT, drug therapy  
 sulfamethoxazole: CB, drug combination  
 sulfamethoxazole: DT, drug therapy  
 trimethoprim: CB, drug combination  
 trimethoprim: DT, drug therapy  
 cotrimoxazole: DT, drug therapy  
 hyaluronic acid: DT, drug therapy  
 povidone iodine: DT, drug therapy  
 sulfadiazine silver: DT, drug therapy  
 framycetin: DT, drug therapy  
 tetanus toxoid: DT, drug therapy  
 midazolam: DT, drug therapy  
 cefalexin: DT, drug therapy  
 gentamicin: DT, drug therapy  
 propanol: DT, drug therapy  
 2 propanol: DT, drug therapy  
 biphenyl derivative: DT, drug therapy  
 ringer lactate solution: DT, drug therapy  
 hydrogen peroxide: AE, adverse drug reaction  
 hydrogen peroxide: DT, drug therapy  
 drug delivery system: DT, drug therapy  
 connettivina  
 tetanol  
 te anatoxal  
 midazolam maleate  
**gentamicin bone cement**  
 \*local anesthetic agent: AE, adverse drug reaction  
 \*local anesthetic agent: CM, drug comparison  
 \*local anesthetic agent: DT, drug therapy  
 ornipressin: DT, drug therapy  
 pethidine: DT, drug therapy  
 sofratulle  
 unclassified drug  
 kodan  
 oracet  
 ialugen  
 rinkilast

L74 ANSWER 26 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.  
 93230115 EMBASE Evidence of hyaluronic acid and hyaluronic acid binding sites on human corneal endothelium. Harfstrand A.; Molander N.; Stenevi U.; Apple D.; Schenholm M.; Madsen K.. Kabi Pharmacia Ophthalmics AB, S-751 82 Uppsala, Sweden. J. CATARACT REFRACTIVE SURG. 18/3 (265-269) 1992. ISSN: 0886-3350. CODEN: JCSUEV. Pub. Country: United States. Language: English. Summary Language: English.

AB A highly specific hyaluronic acid (HA) recognizing protein (HABR) was used to study whether the human corneal endothelium is covered by HA and to quantify the amount. Tritiated high molecular weight HA was used to determine the capacity of the human endothelium to bind exogenous HA. Human corneas were obtained from keratoconus patients having corneal transplantation and from postmortem eyes. The corneas



were immersed in a 4% formaldehyde solution containing 1% cetylpyridine chloride for histochemistry, frozen for biochemistry, or used for 3H-HA (M(r) 3 x 106) binding. For the biochemical determinations, 125I-labeled HABR was used. Tritiated HA was used for the binding experiment. A specific layer of HA covering the endothelial cells of the corneal buttons was demonstrated. The biochemical analysis also revealed the presence of HA. Finally, the human endothelial cells had specific hyaluronic acid binding sites.

CT EMTAGS: visual system (0915); prevention (0165); histology (0330); rabbits and hares (0731); mammal (0738); human (0888); nonhuman (0777); human tissue, cells or cell components (0111); animal tissue, cells or cell components (0105); article (0060)

Medical Descriptors:

\*binding site  
 \*protein binding  
 \*cornea endothelium  
 binding affinity  
 keratoconus  
 cell protection  
**cornea transplantation**  
 molecular weight  
 surgical technique  
 histochemistry  
 aqueous humor  
 cell migration  
 rabbit  
 antibody specificity  
 human  
 nonhuman  
 human cell  
 animal cell  
 article  
 Drug Descriptors:  
 \*hyaluronic acid  
 formaldehyde  
 cetylpyridinium salt  
 glycosaminoglycan

L74 ANSWER 27 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.

92149166 EMBASE Cemented versus cementless hip arthroplasty A review of prosthetic biocompatibility. Santavirta S.; Gristina A.; Konttinen Y.T.. Orthopedic Hospital of the, Invalid Foundation, Tenholantie 10, SF-00280 Helsinki, Finland. ACTA ORTHOP. SCAND. 63/2 (225-232) 1992. ISSN: 0001-6470. CODEN: AOSAAK. Pub. Country: Denmark. Language: English. Summary Language: English.

AB The fibrous interface tissue between hip prostheses and surrounding bone is often morphologically and functionally synovial-like. The fibroblast is the major cell type; but also giant cells and macrophages are present, and their numbers are increased in the occasional adverse-type host reaction to the prosthesis. Adverse lytic reactions are often associated with methylmethacrylate debris, whereas in cementless cases, polyethylene and metallic (titanium) wear debris seem to cause adverse reactions. Osteoblasts, osteoclasts, and mesenchymal collagenase secreted by fibroblasts and macrophages play an important role in the process of prosthetic loosening. Methylmethacrylate is immunologically relatively inert, while it induces inflammatory mononuclear-cell migration. Both cemented and cementless prostheses cause a foreign-body type host response, including adaptive and reactive processes. This response includes the formation of fibroblast-like B-type lining cells, which are able to synthesize and secrete hyaluronate. Material surfaces of hip arthroplasty components also provide a unique environmental niche to which staphylococcal strains adhere and colonize. Antibiotic resistance is related to the material colonized rather than to the presence of an exopolysaccharide barrier, organisms bound to polyethylene and methylmethacrylate are more resistant than organisms that are bound to stainless steel. An understanding of prosthetic biocompatibility requires an appreciation of tissue cell, bacterial cell and host defense-system response to biomaterials. The

site of implantation is a stage on which the 'players' (bacteria, host cells, and organic moieties) interact and compete, and before which the host is a 'responsive audience.'

CT EMTAGS: apparatus, equipment and supplies (0510); reticuloendothelial system (0924); etiology (0135); bacterium (0762); **infection** (0310); priority journal (0007); review (0001); enzyme (0990)

Medical Descriptors:

\*hip arthroplasty

**\*biocompatibility**

\*hip prosthesis

fibroblast

giant cell

macrophage

prosthesis loosening

osteoblast

osteoclast

mononuclear cell

foreign body reaction: ET, etiology

bacterium adherence

staphylococcus

antibiotic resistance

immunopathology

immune response

**bacterial infection**

priority journal

review

Drug Descriptors:

cement

methacrylic acid methyl ester

polyethylene

titanium

collagenase: EC, endogenous compound

**hyaluronic acid: EC, endogenous compound**

L74 ANSWER 28 OF 51 MEDLINE

92252289 [Comparative studies of the use of viscoelastic substances in cataract surgery. A randomized study]. Vergleichende Untersuchungen zum Einsatz von visko-elastischen Substanzen in der Kataraktchirurgie. Eine randomisierte Studie. Ozmen A; Guthoff R; Winter R; Draeger J. (Universitäts-Augenklinik Hamburg.. )KLINISCHE MONATSBLÄTTER FÜR AUGENHEILKUNDE, (1992 Mar) 200 (3) 171-4. Journal code: KWA. ISSN: 0023-2165. Pub. country: GERMANY: Germany, Federal Republic of. Language: German.

AB In three prospectively randomized groups of patients viscoelastic materials during IOL-implantation have been compared concerning 1. intraocular pressure, 2. endothelial cell count, 3. corneal thickness. Examinations were performed preoperatively, the first, second and fifth postoperative day. There was no statistical difference between hydroxypropylmethylcellulose (2%), hyaluronic acid (1%) and air. Examinations were performed preoperatively the first, the second and the fifth postoperative day. There was no statistically significant difference between all groups of patients. Advantages and disadvantages for routine use of viscoelastic substances are discussed.

CT Check Tags: Comparative Study; Human

Corneal Stroma: DE, drug effects

Endothelium, Corneal: DE, drug effects

English Abstract

**Foreign-Body Reaction: ET, etiology**

**\*Hyaluronic Acid: AD, administration & dosage**

Intraocular Pressure: DE, drug effects

**\*Lenses, Intraocular**

**\*Methylcellulose: AA, analogs & derivatives**

Methylcellulose: AD, administration & dosage

**\*Postoperative Complications: ET, etiology**

Prospective Studies

L74 ANSWER 29 OF 51 MEDLINE

93127396 [Current methods of surgical rehabilitation in traumatic retinal detachment]. Sovremennye metody khirurgicheskoi reabilitatsii pri travmaticheskoi otsloike setchatki. Morozova I V; Kiseleva O A. VESTNIK OFTALMOLOGII, (1992 May-Jun) 108 (3) 41-5. Ref: 103. Journal code: XAO. ISSN: 0042-465X. Pub. country: RUSSIA: Russian Federation. Language: Russian.

CT Check Tags: Comparative Study; Human

\*Eye Injuries: CO, complications

\*Hyaluronic Acid

\*Implants, Artificial

Retinal Detachment: ET, etiology

Retinal Detachment: RH, rehabilitation

\*Retinal Detachment: SU, surgery

\*Scleroplasty: MT, methods

\*Silicones

\*Vitreous Body: SU, surgery

L74 ANSWER 30 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.

92041511 EMBASE Low-molecular-weight sodium hyaluronate in the treatment of bacterial corneal ulcers. Gandolfi S.A.; Massari A.; Orsoni J.G.. Istituto di Oftalmologia, Universita di Parma, Via Gramsci 14, I-43100 Parma, Italy. GRAEFE'S ARCH. CLIN. EXP. OPHTHALMOL. 230/1 (20-23) 1992. ISSN: 0721-832X. CODEN: GACODL. Pub. Country: Germany, Federal Republic of. Language: English. Summary Language: English.

AB A double-blind clinical trial was performed on 26 patients suffering from corneal ulcers of proven (i.e., culture-positive) bacterial etiology. After their recruitment, the subjects were randomly assigned to one of the following treatment protocols: (1) tobramycin (15 mg/ml) in saline applied at 1 drop/h or (2) tobramycin (15 mg/ml) in low-molecular-weight hyaluronic acid applied at 1 drop/h. The sample size was adjusted according to a type I error of 0.01 and type a II error of 0.05 for a minimal expected difference of 35%. The healing time was calculated from the beginning of treatment to the day on which a follow-up fluorescein test proved to be negative. The mean healing time (.-.SD) was 3.5 .+- 0.9 days in the sodium hyaluronate group and 5.9 .+- 1.5 days in the saline group (P < 0.001). These results suggest that treatment with an antibiotic dissolved in low-molecular-weight sodium hyaluronate can further shorten the clinical course of a bacterial corneal ulcer.

CT EMTAGS: therapy (0160); infection (0310); mammal (0738); human (0888); male (0041); female (0042); clinical article (0152); adolescent (0017); aged (0019); adult (0018); topical drug administration (0186); priority journal (0007); conference paper (0061); human experiment (0104)

Medical Descriptors:

\*cornea ulcer: DT, drug therapy

\*bacterial infection: DT, drug therapy

\*ulcer healing

viscosity

disease duration

human

male

female

clinical article

adolescent

aged

adult

topical drug administration

priority journal

conference paper

Drug Descriptors:

\*hyaluronic acid: CT, clinical trial

\*hyaluronic acid: CB, drug combination

\*hyaluronic acid: DT, drug therapy

\*tobramycin: CT, clinical trial

\*tobramycin: CB, drug combination

\*tobramycin: DT, drug therapy

\*eye drops: DT, drug therapy

antibiotic agent: CM, drug comparison  
antibiotic agent: DT, drug therapy  
fluorescein

## L74 ANSWER 31 OF 51 MEDLINE

92333799 Effects of residual sodium hyaluronate on postsurgical blood-aqueous barrier. Tsurimaki Y; Shimizu H. (Department of Ophthalmology, Jichi Medical School, Tochigi, Japan.. ) JAPANESE JOURNAL OF OPHTHALMOLOGY, (1991) 35 (4) 446-52. Journal code: KN1. ISSN: 0021-5155. Pub. country: Japan. Language: English.

AB The effects of residual sodium hyaluronate (HA) on the postsurgical blood-aqueous barrier (BAB) function were investigated in 79 posterior chamber lens (PCL)-implanted eyes after both extracapsular cataract extraction and PCL implantation using HA products. The amount of residual HA was classified according to the status of the aqueous warm current on the 1st postoperative day. The eyes with static warm current were classified into the static current group and the other eyes into the normal current group. Aqueous flare intensity and cell number were measured in all eyes daily from the 1st to the 7th postoperative day using the flare-cell meter. Of the 79 eyes, 11 eyes (14%) were classified into the static current group. Flare intensity showed the most marked difference between the two groups on the 1st postoperative day. The difference was statistically significant from the 1st to the 7th postoperative days (P less than 0.05). Cell count was also higher in the static current group throughout the observation period except for the 3rd and 4th postoperative days (P less than 0.05). These findings suggest that residual HA exacerbated the postoperative inflammation and that its effects on the BAB continued for at least a week.

CT Check Tags: Female; Human; Male

Aged

\*Aqueous Humor: ME, metabolism

Biological Transport, Active

\*Blood: ME, metabolism

\*Cataract Extraction

Cell Count

Endophthalmitis: ET, etiology

\*Hyaluronic Acid: PK, pharmacokinetics

\*Lenses, Intraocular

Postoperative Complications

## L74 ANSWER 32 OF 51 MEDLINE

91348970 Hyaluronic acid prevents oxygen free-radical damage to granulation tissue: a study in rats. Foschi D; Castoldi L; Radaelli E; Abelli P; Calderini G; Rastrelli A; Mariscotti C; Marazzi M; Trabucchi E. (Department of Surgery, Institute of Biomedical Sciences L. Sacco, Milan, Italy.. ) INTERNATIONAL JOURNAL OF TISSUE REACTIONS, (1990) 12 (6) 333-9. Journal code: GTG. ISSN: 0250-0868. Pub. country: Switzerland. Language: English.

AB Oxygen free-radicals are known to impair wound healing after ischaemia-reperfusion or polymorphonuclear cell stimulation. Furthermore, they reduce the breaking strength of all recent wounds and might be a cause of wound leakage. This study was performed to evaluate whether or not hyaluronic acid can reduce the risk of wound impairment caused by free-radicals, in rats with abdominal sepsis, polymorphonuclear cell stimulation or cytochrome C function derangement produced by xenobiotics. Male Sprague-Dawley rats with open wounds received phenazine methosulfate or zimosan, or had abdominal sepsis to induce oxygen free-radical generation. There were three groups of treatment: hyaluronic acid cream, hyaluronic acid ethyl ester gel, and placebo. The reduction in wound size was measured from the 1st to the 11th postoperative day; biopsies were taken for histological evaluation. Every other day, a gentle debridement was performed in all the groups of animals. We found that hyaluronic acid and its ethyl ester derivative significantly improved the wound healing of rats subjected to an increased generation of oxygen free-radicals. It remains to be established whether or not hyaluronic acid acts as a scavenger of free-radicals.

CT Check Tags: Animal; Male

**Bacterial Infections: PP, physiopathology**

Cecum: IN, injuries

Cytochrome c: PH, physiology

Free Radicals

\*Granulation Tissue: DE, drug effects

Granulation Tissue: PA, pathology

Granulation Tissue: PP, physiopathology

\*Hyaluronic Acid: PD, pharmacology

**Hyaluronic Acid: TU, therapeutic use**

Methylphenazonium Methosulfate: PD, pharmacology

Neutrophils: DE, drug effects

Neutrophils: PH, physiology

Oxygen: ME, metabolism

\*Oxygen: PD, pharmacology

Rats

Rats, Inbred Strains

Wound Healing: DE, drug effects

Wound Healing: PH, physiology

**Wounds, Penetrating: DT, drug therapy**

Zymosan: PD, pharmacology

L74 ANSWER 33 OF 51 MEDLINE

89041006 Comparison of Healon and Viscoat in cataract extraction and intraocular lens implantation. Alpar J J; Alpar A J; Baca J; Chapman D. (Saint Luke Eye Institute, Panhandle Ophthalmological Foundation, Amarillo, Texas 79106-4161.. )OPHTHALMIC SURGERY, (1988 Sep) 19 (9) 636-42. Journal code: OIC. ISSN: 0022-023X. Pub. country: United States. Language: English.

AB Sixty patients were randomly assigned to Healon (20 patients) or Viscoat (40 patients) treatment during extracapsular cataract extraction and intraocular lens implantation surgery. The 40 patients in the Viscoat group were randomly subdivided into two groups. In one group (20 patients), Viscoat was irrigated/aspirated from the eye at the close of surgery, while in the second group of 20 patients, Viscoat was left in the eye. In all Healon cases, the viscoelastic substance was removed from the eye at the end of the surgical procedure. Compared with Viscoat, Healon better facilitated the surgical procedure and appeared to be a more advantageous viscoelastic preparation. Viscoat, in many cases, caused rises in intraocular pressure in the immediate postoperative period when either removed or left in the eye at the close of surgery.

CT Check Tags: Comparative Study; Human

Cataract Extraction: AE, adverse effects

\*Cataract Extraction: MT, methods

Cell Count

\*Chondroitin: TU, therapeutic use

Cornea: PA, pathology

Drug Combinations: TU, therapeutic use

Drug Evaluation

**Endophthalmitis: ET, etiology****Endophthalmitis: PA, pathology**

Endothelium, Corneal: PA, pathology

\*Hyaluronic Acid: TU, therapeutic use

Intraocular Pressure: DE, drug effects

**Lenses, Intraocular: AE, adverse effects**

\*Lenses, Intraocular: MT, methods

Postoperative Period

Time Factors

L74 ANSWER 34 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.

86217952 EMBASE Diffuse cutaneous hypersensitivity reaction after dexamethasone/polymyxin B/neomycin combination eyedrops. Baldinger J.; Weiter J.J.. Eye Research Institute of Retina Foundation, Boston, MA 02114, United States. ANN. OPHTHALMOL. 18/3 (95-96) 1986. CODEN: ANOPB5. Pub. Country: United States. Language: English.

AB Localized cutaneous hypersensitivity reactions to antibiotic eyedrops are not unusual. To our knowledge, however, a diffuse cutaneous reaction to eyedrops containing dexamethasone/polymyxin B/neomycin has never been reported. We describe the diffuse skin

changes noted in a 72-year-old patient five days after starting eyedrop therapy.

CT EMTAGS: priority journal (0007); immunological factors (0136); skin, hair, nails and sweat glands (0980); therapy (0160); adverse drug reaction (0198); intoxication (0302); topical drug administration (0186); case report (0151); human (0888); visual system (0915)

Medical Descriptors:

\*pharmacotherapy  
\*adverse drug reaction  
\*drug hypersensitivity  
\*skin toxicity  
\*dexamethasone  
\*polymyxin b  
\*neomycin  
\*skin allergy  
\*eye drops  
\*skin  
\*hypersensitivity  
gentamicin  
hyaluronic acid  
acetylcholine  
cyclopentolate

L74 ANSWER 35 OF 51 MEDLINE

86171308 Management of a posterior capsule rupture in planned extracapsular cataract extraction and posterior chamber lens implantation. Wang H S. JOURNAL OF CATARACT AND REFRACTIVE SURGERY, (1986 Jan) 12 (1) 73-6. Journal code: JPB. ISSN: 0886-3350. Pub. country: United States. Language: English.

AB Management of zonular dialysis and posterior capsule rupture during extracapsular cataract extraction is described. The Heslin gravity cannula is advocated to maintain the normal structure of the anterior segment in a closed chamber technique. Lens cortical material is stripped away using the manual technique described by Gills and McIntyre to avoid vitreous loss. It is then possible to proceed with posterior chamber lens implantation. If vitreous loss occurs, an adequate anterior vitrectomy with an automated vitreous cutter is recommended. A posterior chamber lens implant is preferred if there is adequate capsule to support the lens.

CT Check Tags: Human  
Cataract Extraction: IS, instrumentation  
\*Cataract Extraction: MT, methods  
**Hyaluronic Acid: AD, administration & dosage**  
\*Intraoperative Complications: SU, surgery  
\*Lens Capsule, Crystalline: IN, injuries  
Lens Capsule, Crystalline: SU, surgery  
\*Lens, Crystalline: IN, injuries  
**\*Lenses, Intraocular Rupture**  
Suture Techniques: IS, instrumentation  
Vitrectomy: IS, instrumentation  
Vitrectomy: MT, methods

L74 ANSWER 36 OF 51 MEDLINE

86106527 [Use of Healon in surgery of the anterior segment. Apropos of 52 cases]. De l'utilisation du Healon dans la chirurgie du segment anterieur. A propos de 52 cas. Lagoutte F; Di Battista J C; Banos M T. BULLETIN DES SOCIETES D OPHTALMOLOGIE DE FRANCE, (1985 Feb) 85 (2) 277-8. Journal code: C40. ISSN: 0081-1270. Pub. country: France. Language: French.

CT Check Tags: Human  
Adult  
Anterior Chamber  
\*Anterior Eye Segment: SU, surgery  
Child  
**Cornea: TR, transplantation**  
**Corneal Transplantation**  
**Eye Injuries: SU, surgery**

**\*Hyaluronic Acid: AD, administration & dosage**  
**Injections**  
**Lenses, Intraocular**

L74 ANSWER 37 OF 51 MEDLINE

85029726 Viscous corneal protection by sodium hyaluronate, chondroitin sulfate, and methylcellulose. Hammer M E; Burch T G. INVESTIGATIVE OPHTHALMOLOGY AND VISUAL SCIENCE, (1984 Nov) 25 (11) 1329-32. Journal code: GWI. ISSN: 0146-0404. Pub. country: United States. Language: English.

AB The authors' study of the viscosities of various concentrations of sodium hyaluronate, chondroitin sulfate, and methylcellulose revealed that sodium hyaluronate and methylcellulose are pseudoplastic fluids in contrast to chondroitin sulfate, which is a Newtonian fluid. Pseudoplastic fluids are ideal for maintaining the anterior chamber, since they are more viscous at rest. Intermediate viscosity preparations of these three agents used as a thin endothelial coating gave excellent protection from intraocular lens abrasion. A highly viscous agent, eg, sodium hyaluronate 1%, in a thin layer produced extensive endothelial cell damage because it transmitted excessive shear force to the endothelium. A highly viscous agent, sodium hyaluronate 1% in a thick layer produced a physical barrier to compression with little endothelial damage. A low-viscosity agent, balanced salt solution provided insufficient protection against intraocular lens abrasion.

CT Check Tags: Animal; Comparative Study  
 Cattle  
 Chemistry  
 \*Chondroitin: AA, analogs & derivatives  
 \*Chondroitin Sulfates: TU, therapeutic use  
 \*Cornea  
**Eye Injuries: PC, prevention & control**  
**\*Hyaluronic Acid: TU, therapeutic use**  
 Hydrogen-Ion Concentration  
**Lenses, Intraocular: AE, adverse effects**  
 \*Methylcellulose: TU, therapeutic use  
 Viscosity

L74 ANSWER 38 OF 51 MEDLINE

84143487 Combined use of sodium hyaluronate and tissue adhesive in penetrating keratoplasty of corneal perforations. Maguen E; Nesburn A B; Macy J I. OPHTHALMIC SURGERY, (1984 Jan) 15 (1) 55-7. Journal code: OIC. ISSN: 0022-023X. Pub. country: United States. Language: English.

AB A new technique which allows the use of a guarded trephine in penetrating keratoplasty for corneal perforation is described. It involves the combined use of cyanoacrylate adhesive and sodium hyaluronate and allows a normotensive eye to be obtained prior to trephination. Five cases in which this technique was used are described. A better tectonic result can be obtained and visual rehabilitation may be more readily achieved without performing a secondary procedure.

CT Check Tags: Case Report; Female; Human; Male; Support, Non-U.S. Gov't; Support, U.S. Gov't, P.H.S.  
 Aged  
**Bacterial Infections: CO, complications**  
**\*Cornea: TR, transplantation**  
 Corneal Diseases: CO, complications  
 \*Corneal Diseases: SU, surgery  
**\*Corneal Transplantation**  
**\*Corneal Ulcer: SU, surgery**  
 Cyanoacrylates: TU, therapeutic use  
**\*Hyaluronic Acid: TU, therapeutic use**  
 Keratitis, Dendritic: CO, complications  
 Middle Age  
 Surgical Instruments  
 \*Tissue Adhesives: TU, therapeutic use

L74 ANSWER 39 OF 51 MEDLINE

83242682 [Healon as an emergency aid]. Healon als Nothelfer. Neubauer H. KLINISCHE MONATSBLETT FÜR AUGENHEILKUNDE, (1983 Apr) 182 (4) 269-71. Journal code: KWA. ISSN: 0023-2165. Pub. country: GERMANY, WEST: Germany, Federal Republic of. Language: German.

AB Sodium hyaluronate (Healon) was used in follow-up surgery of the anterior segment. In addition to the uses of Healon already documented in the literature the author points out that very early synechiolysis with it can be beneficial especially in cases of distortion of the pupil after cataract extraction with insufficient cleaning of the vitreous. So far Healon has not caused any problems with regard to tissue compatibility, viscosity, resorption or its optical quality in the anterior chamber. The examples of secondary treatment given after injuries are intended to encourage the use of Healon in risky situations as a means of stabilizing the anterior chamber with a viscous fluid.

CT Check Tags: Case Report; Female; Human; Male

Adult

Anterior Chamber: DE, drug effects

Cataract Extraction: AE, adverse effects

Drug Evaluation

\*Emergencies

English Abstract

**Eye Foreign Bodies: DT, drug therapy**

**Eye Injuries: DT, drug therapy**

\***Hyaluronic Acid: TU, therapeutic use**

**Lenses, Intraocular: AE, adverse effects**

Postoperative Care

Vitreous Body: DE, drug effects

L74 ANSWER 40 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.

84080391 EMBASE Visco elastic materials in keratoplasty. Steele A.D.McG.. Moorfields Eye Hospital, London EC1V 2PD, United Kingdom. TRANS. OPHTHALMOL. SOC. U. K. 103/3 (268-269) 1983. CODEN: TOSUAH. Pub. Country: United Kingdom. Language: English.

CT EMTAGS: drug comparison (0196); visual system (0915); topical drug administration (0186); clinical article (0152); therapy (0160); human (0888)

Medical Descriptors:

\*drug comparison

\***cornea transplantation**

\***cornea graft**

\*donor

\***hyaluronic acid**

\*hydroxypropylmethylcellulose

betamethasone

**gentamicin**

L74 ANSWER 41 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.

84112107 EMBASE Bacterial keratitis following radial keratotomy. Wilhelmus K.R.; Hamburg S.. Cullen Eye Institute, Baylor College of Medicine, Houston, TX 77030, United States. CORNEA 2/2 (143-146) 1983. CODEN: CORNDB. Pub. Country: United States. Language: English.

CT EMTAGS: therapy (0160); topical drug administration (0186); human (0888); visual system (0915); **infection** (0310); case report (0151); bacterium (0762); skin, hair, nails and sweat glands (0980)

Medical Descriptors:

\*pharmacotherapy

\*staphylococcus aureus

\***refractive keratoplasty**

\*keratotomy

\*keratitis

\*myopia

\***penicillin g**

\***metacillin**

\***ampicillin**

\***cefalotin**

\***vancomycin**

\***erythromycin**



- \*chloramphenicol
- \*tetracycline
- \*clindamycin
- \*kanamycin
- \*gentamicin
- \*cefazolin
- \*prednisolone
- \*hyaluronic acid
- refraction

L74 ANSWER 42 OF 51 MEDLINE

84153691 Anterior segment viscosurgery with Healon. Proceedings of Australian seminars, May 1983. Anonymous. AUSTRALIAN JOURNAL OF OPHTHALMOLOGY, (1983 Aug) 11 (3 Suppl) 1-26. Journal code: 9G5. ISSN: 0310-1177. Pub. country: Australia. Language: English.

CT Check Tags: Human

- \*Anterior Chamber: SU, surgery
- Cataract Extraction: IS, instrumentation
- \*Eye: SU, surgery
- Eye Injuries: SU, surgery**
- \*Hyaluronic Acid: TU, therapeutic use**
- Lenses, Intraocular**

L74 ANSWER 43 OF 51 MEDLINE

83030499 Assessment of intraocular lens implantation in children. Menezo J L; Taboada J. JOURNAL - AMERICAN INTRA-OCULAR IMPLANT SOCIETY, (1982 Spring) 8 (2) 131-5. Journal code: HA1. ISSN: 0146-2776. Pub. country: United States. Language: English.

AB Visual rehabilitation by conventional aphakic spectacles and contact lenses has posed a serious problem in the pediatric population. While the use of intraocular lenses has not achieved widespread acceptance as a form of aphakic correction, we have obtained encouraging results in some of our patients. In this report, we discuss two categories: those patients with congenital cataracts, and those patients with traumatic cataracts.

CT Check Tags: Human

- Age Factors
- Amblyopia: PC, prevention & control
- \*Aphakia, Postcataract: RH, rehabilitation
- \*Cataract: CN, congenital
- Cataract: ET, etiology
- Cataract Extraction: MT, methods
- Child
- Eye Injuries: CO, complications**
- Hyaluronic Acid: TU, therapeutic use**
- Intraoperative Complications
- Lens Capsule, Crystalline: SU, surgery
- \*Lenses, Intraocular**
- Postoperative Complications

L74 ANSWER 44 OF 51 EMBASE COPYRIGHT 1996 ELSEVIER SCI. B.V.

81253290 EMBASE New drugs in ophthalmology. Taylor R.F.. Roy. Prince Alfred Hosp., Sydney, Australia. AUST. J. OPHTHAL. 9/3 (246-248) 1981. CODEN: AJOHBL. Pub. Country: Australia. Language: English.

CT EMTAGS: heart (0921); respiratory system (0930); adverse drug reaction (0198); visual system (0915); **infection** (0310); short survey (0002); topical drug administration (0186)

Medical Descriptors:

- \*hyaluronic acid**
- \*healon
- \*timolol
- \*intraocular hypertension
- \*superficial punctate keratitis
- \*anxiety
- \*bradycardia
- \*bronchospasm
- \*adverse drug reaction
- \*cornea transplantation**
- \*cataract extraction

- \*eye surgery
- \*virus infection
- \*trifluridine
- \*aciclovir
- \*dipivefrine
- \*gliclazide

## L74 ANSWER 45 OF 51 MEDLINE

82257833 [Use of a viscous substance (Healon) in spatial tactics of the anterior segment]. L'emploi d'une substance visqueuse (Healon) dans la tactique spatiale du segment anterieur. Eisner G. BULLETINS ET MEMOIRES DE LA SOCIETE FRANCAISE D OPHTALMOLOGIE, (1981) 93 201-6. Journal code: BP4. ISSN: 0081-1092. Pub. country: France. Language: French.

CT Check Tags: Human

\*Anterior Chamber: SU, surgery

**Eye Injuries: SU, surgery**

\*Hyaluronic Acid: TU, therapeutic use

**Lenses, Intraocular**

Methods

## L74 ANSWER 46 OF 51 MEDLINE

81152818 Hyaluronic acid stimulates neutrophil function in vitro and in vivo. A review of experimental results and a presentation of a preliminary clinical trial. Hakansson L; Hallgren R; Venge P; Artursson G; Vedung S. SCANDINAVIAN JOURNAL OF INFECTIOUS DISEASES. SUPPLEMENTUM, (1980) Suppl 24 54-7. Journal code: UCY. ISSN: 0300-8878. Pub. country: Sweden. Language: English.

AB Hyaluronic acid (HA) stimulates normal neutrophil function both in vitro and in vivo Stimulation was also achieved by subcutaneous administration of HA to patients with extreme susceptibility to bacterial infections. Clinical improvement of some patients was obtained in connection to the administration. It is premature at this time to conclude any therapeutic effect of HA in patients with extreme infection propensity. The data presented here, however, for certain merit further investigation on this matter.

CT Check Tags: Female; Human; Male; Support, Non-U.S. Gov't

Adult

Aged

**Bacterial Infections: DT, drug therapy****Burns: DT, drug therapy**

Cells, Cultured

Child

Clinical Trials

\*Hyaluronic Acid: PD, pharmacology

**Hyaluronic Acid: TU, therapeutic use**

Middle Age

\*Neutrophils: DE, drug effects

Neutrophils: IM, immunology

Neutrophils: ME, metabolism

Phagocytosis: DE, drug effects

## L74 ANSWER 47 OF 51 MEDLINE

72256737 Nature of the bond between partial-thickness skin and wound granulations. Burleson R; Eiseman B. SURGERY, (1972 Aug) 72 (2) 315-22. Journal code: VC3. ISSN: 0039-6060. Pub. country: United States. Language: English.

CT Check Tags: Animal

Collagen: PH, physiology

Elastin: PH, physiology

\*Granulation Tissue

Heparin: PD, pharmacology

**Hyaluronic Acid: PH, physiology**

Hyaluronidase: PD, pharmacology

Microbial Collagenase: PD, pharmacology

Pancreatopeptidase: PD, pharmacology

Plasmin: PD, pharmacology

Rats

\*Skin: TR, transplantation

rcu  
3-18

**\*Skin Transplantation**

Swine

Thrombin: PD, pharmacology

**Transplantation, Autologous****Transplantation, Heterologous**

Trypsin: PD, pharmacology

Wound Healing: DE, drug effects

**Wound Infection: PP, physiopathology**

L74 ANSWER 48 OF 51 MEDLINE

71291136 Chemical and osmolar changes of interstitial fluid in acute inflammatory states. Vakili C; Ruiz-Ortiz F; Burke J F. SURGICAL FORUM, (1970) 21 227-8. Journal code: VB0. ISSN: 0071-8041. Pub. country: United States. Language: English.

CT Check Tags: Animal

Ascorbic Acid: PD, pharmacology

**\*Extracellular Space: AN, analysis**

Hexosamines: AN, analysis

**Hyaluronic Acid: PD, pharmacology**

Inflammation: CI, chemically induced

Inflammation: ET, etiology

**\*Inflammation: ME, metabolism****\*Osmolar Concentration**

Potassium: AN, analysis

Proteins: AN, analysis

Sodium: AN, analysis

**Staphylococcal Infections: CO, complications**

Uronic Acids: AN, analysis

**Wounds and Injuries: CO, complications**

L74 ANSWER 49 OF 51 MEDLINE

70107784 [The intermediate substance of the aorta in dissecting aneurysm in comparison with various diseases]. Untersuchung uber die Intermediarsubstanz der Aorta bei Aneurysma dissecans im Vergleich zu anderen Krankheitsbildern. Jozsa L; Szederkenyi G; Lusztig G. ACTA BIOLOGICA ET MEDICA GERMANICA, (1969) 23 (2) 323-8. Journal code: OE6. Pub. country: GERMANY, EAST: German Democratic Republic. Language: German.

CT Check Tags: Female; Human; Male

**\*Aorta: AN, analysis**

Aortic Aneurysm: CO, complications

**\*Aortic Aneurysm: ME, metabolism****\*Aortic Diseases: ME, metabolism****Aortic Rupture: ET, etiology**

Arteriosclerosis: ME, metabolism

Autopsy

Cardiac Tamponade: ET, etiology

**\*Chondroitin: AN, analysis**

Cushing's Syndrome: ME, metabolism

Diabetic Angiopathies: ME, metabolism

**\*Glycosaminoglycans: AN, analysis****\*Heparin: AN, analysis****\*Hyaluronic Acid: AN, analysis**

Hyperthyroidism: ME, metabolism

Hypothyroidism: ME, metabolism

Middle Age

Sulfates: AN, analysis

**Syphilis, Cardiovascular: ME, metabolism**

L74 ANSWER 50 OF 51 MEDLINE

71153630 [Problems in vitreous body surgery and indications for intravitreous injections]. Les difficultes de la chirurgie vitreenne et les indications des injections intra-vitreennes. Moreau P G; Pichon P. BULLETINS ET MEMOIRES DE LA SOCIETE FRANCAISE D OPHTALMOLOGIE, (1968) 81 74-9. Journal code: BP4. ISSN: 0081-1092. Pub. country: France. Language: French.

CT Check Tags: Human

Cataract: ET, etiology

Cerebrospinal Fluid

**Eye Foreign Bodies: SU, surgery**  
**Eye Injuries: SU, surgery**  
 Freeze Drying  
 Glaucoma: ET, etiology  
 Hemorrhage: ET, etiology  
**Hyaluronic Acid: TU, therapeutic use**  
**Infection: ET, etiology**  
 \*Injections  
   Methods  
   Postoperative Complications  
   Retinal Detachment: SU, surgery  
   Silicones: TU, therapeutic use  
   Tissue Therapy  
 \*Vitreous Body: SU, surgery

L74 ANSWER 51 OF 51 MEDLINE

70250123 [The effect of hyaluronidase, hyaluronic acid and several other substances on post-radiation experimental bacteremia]. Vliianie gialuronidazy, gialuronovoi kisloty i nekotorykh drugikh veshchestv na postradiatsionnuu eksperimental'nuiu bakteriemiiu. Alaverdian M I; Ter-Avetisian A T. BIULLETEN EKSPERIMENTALNOI BIOLOGII I MEDITSINY, (1967 Sep) 64 (9) 51-3. Journal code: A74. ISSN: 0006-4041. Pub. country: USSR. Language: Russian.

CT Check Tags: Animal  
   Chlortetracycline: TU, therapeutic use  
   English Abstract  
   Epinephrine: TU, therapeutic use  
 \*Hyaluronic Acid: TU, therapeutic use  
 \*Hyaluronidase: PD, pharmacology  
   Mice  
   Rabbits  
 \*Radiation Injuries, Experimental: CO, complications  
   Radiation Injuries, Experimental: DT, drug therapy  
   Radiation Injuries, Experimental: ET, etiology  
   Septicemia: CO, complications  
 \*Septicemia: DT, drug therapy  
   Vitamin K: TU, therapeutic use

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E FALLK R/AU  
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 L75 25 S E3,E6  
   E ASCULAI S/AU  
 L76 20 S E4  
 L77 14 S L75 AND L76  
 L78 805 S HYALURONIC  
 L79 51 S ?INFECT? AND L78  
 L80 65 S ?TOPICAL? AND L78  
 L81 13 S L79 AND L80  
 L82 63 S (?IMPLANT? OR ?TRANSPLANT?) AND L78  
 L83 3 S L82 AND L79  
 L84 4 S L82 AND L80

L85 20 S L81 OR L83 OR L84  
L86 33 S L85 OR L77

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=> d 1-33 bib abs

L86 ANSWER 1 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD

AN 96-030916 [04] WPIDS

DNC C96-010656

TI Use of hyaluronic acid - to prevent arterial restenosis after balloon angioplasty and narrowing of tubular walls after traumatisation.

DC B04

IN **ASCULAI, S S; FALK, R E; TURLEY, E A**

PA (NORP-N) NORPHARMC O INC

CYC 1

PI CA 2120045 A 950926 (9604)\* 7 pp

ADT CA 2120045 A CA 94-2120045 940325

PRAI CA 94-2120045 940325

AN 96-030916 [04] WPIDS

AB CA 2120045 A UPAB: 960129

Preventing the narrowing of the tubular walls of an animal after the tubular walls have been traumatised, comprises admin. of a non-toxic amt. of hyaluronic acid and/or their salts and/or homologues, analogues, derivs., complexes, esters, fragments, and subunits of hyaluronic acid to the animal to prevent removing of the tubular walls.

The compsn. is used in liquid form or intravenous form. The compsn. is in injectable or intravenous form and the hyaluronic acid is sodium hyaluronate.

USE - Hyaluronic acid is believed to prevent stenosis of the inner diameter of irritated tubular walls and partic. prevent restenosis of the arterial walls by e.g. the proliferation of endothelial cells as a result of irritation arising from balloon angioplasty or other treatment. The methods and compsns. can be used to prevent restenosis and inhibit restenosis e.g. post operatively in peripheral vascular systems.

ADVANTAGE - The method is safe and non-toxic.

Dwg.0/21

L86 ANSWER 2 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD

AN 96-010551 [01] WPIDS

DNC C96-003250

TI Treatment of cancers and redn. of cancer metastases - by admin. of dosage forms which contain e.g. sodium hyaluronate with a mol. wt. below 750000 Daltons.

DC B05

IN **ASCULAI, S S; FALK, R E**

PA (NORP-N) NORPHARMC O INC

CYC 63

PI WO 9530423 A2 951116 (9601)\* EN 271 pp

RW: AT BE CH DE DK ES FR GB GR IE IT KE LU MC MW NL OA PT SD SE  
SZ UG

W: AM AT AU BB BG BR BY CA CH CN CZ DE DK EE ES FI GB GE HU IS  
JP KE KG KP KR KZ LK LR LT LU LV MD MG MN MW MX NO NZ PL PT  
RO RU SD SE SG SI SK TJ TT UA UG US UZ VN

ADT WO 9530423 A2 WO 95-CA259 950428

PRAI CA 94-2122519 940429

AN 96-010551 [01] WPIDS

AB WO 9530423 A UPAB: 960108

Use of effective dosage amts. ((A) and (B)) of pharmaceutical compsns. for (a) treatment of cancer in patients, (b) prevention of metastases in patients suffering from cancer, and (c) delivery of drugs to the lymph system and/or liver is new. Dosage amt. (A) comprises an anticancer drug and/or a drug suitable for use in treatment of cancer. It also comprises hyaluronic acid and/or a salt of this, with a mol. wt. < 750000 Daltons. The components are in sterile water suitable for injection. Dosage amt. (B) comprises (i)

hyaluronic acid and/or a salt of this, with a mol. wt. < 750000 Daltons, (ii) a drug selected from non-steroidal antiinflammatory drugs and/or chemotherapeutic agents and opt. (iii) an antioxidant.

USE - The dosage form combination is esp. useful for treatment of breast cancer and prevention of metastases in patients with breast cancer.

ADVANTAGE - The combination reduces the risk of recurrence of the disease.

Dwg.0/0

L86 ANSWER 3 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD

AN 95-403801 [51] WPIDS

DNC C95-173380

TI Compsn. comprising hyaluronic acid - useful for treating atherosclerosis.

DC B04

IN **ASCULAI, S S; FALK, R E**

PA (NORP-N) NORPHARMCO INC

CYC 63

PI WO 9529683 A1 951109 (9551)\* EN 21 pp

RW: AT BE CH DE DK ES FR GB GR IE IT KE LU MC MW NL OA PT SD SE  
SZ UG

W: AM AT AU BB BG BR BY CA CH CN CZ DE DK EE ES FI GB GE HU IS  
JP KE KG KP KR KZ LK LR LT LU LV MD MG MN MW MX NO NZ PL PT  
RO RU SD SE SG SI SK TJ TT UA UG US UZ VN

ADT WO 9529683 A1 WO 95-CA243 950427

PRAI CA 94-2122551 940429

AN 95-403801 [51] WPIDS

AB WO 9529683 A UPAB: 951221

Clearing atherosclerosis comprises administering a dosage amt. of a compsn. comprising a non-toxic amt. of each of a chelating agent, a non-steroidal antiinflammatory drug (NSAID), an antioxidant and a form of hyaluronic acid selected from hyaluronic acid or its salts, homologues, analogues, derivs., esters, complexes, fragments or subunits.

Also claimed are (i) a compsn. as described above for intravenous admin. and a dosage amt. of the compsn. as described above.

USE - The compsn. can be used to treat arterial diseases, partic. atherosclerosis.

Dwg.0/0

L86 ANSWER 4 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD

AN 95-358333 [46] WPIDS

DNC C95-156690

TI Restenosis prevention by hyaluronic acid and analogues - esp. after balloon angioplasty, opt. in presence antiinflammatory drug and/or free radical scavenger.

DC B04

IN **ASCULAI, S S; FALK, R E; TURLEY, E A**

PA (NORP-N) NORPHARMCO INC

CYC 53

PI WO 9526193 A1 951005 (9546)\* EN 89 pp

RW: AT BE CH DE DK ES FR GB GR IE IT LU MC NL OA PT SE

W: AT AU BB BG BR BY CA CH CN CZ DE DK ES FI GB GE HU JP KG KP  
KR KZ LK LU LV MD MG MN MW NL NO NZ PL PT RO RU SD SE SI SK  
TJ TT UA US UZ VN

AU 9464222 A 951017 (9604)

ADT WO 9526193 A1 WO 94-CA188 940325; AU 9464222 A AU 94-64222 940325,  
WO 94-CA188 940325

FDT AU 9464222 A Based on WO 9526193

PRAI WO 94-CA188 940325

AN 95-358333 [46] WPIDS

AB WO 9526193 A UPAB: 951122

Preventing narrowing of tubular walls in an animal after their traumatisation, comprises admin. of hyaluronic acid (HA) and/or its salts and/or homologues, analogues, derivs. complexes, esters, fragments, and subunits.

USE - The method prevents stenosis of the inner dia. of

irritated tubular walls, and partic. restenosis of the arterial walls due to proliferation of endothelial cells after balloon angioplasty or similar treatment. Opt. the HA is combined with: (i) non-steroidal antiinflammatory drugs (NSAIDs) to reduce inflammation; (ii) free radical scavenger(s) and/or antioxidant(s); and (iii) stenosis and restenosis inhibiting drug(s); all of which may enhance the effect  
Dwg.0/21

L86 ANSWER 5 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD  
AN 95-206899 [27] WPIDS  
DNC C95-095886  
TI High affinity integrin binding peptides - can be used to attach cells to a substrate, inhibit the attachment of osteoclasts to bone, promote wound healing, inhibit angiogenesis, metastasis of tumours and migration of smooth muscle cells.  
DC B04 D22  
IN KOIVUNEN, E; RUOSLAHTI, E  
PA (LJOL-N) LA JOLLA CANCER RES FOUND  
CYC 56  
PI WO 9514714 A1 950601 (9527)\* EN 86 pp  
RW: AT BE CH DE DK ES FR GB GR IE IT KE LU MC MW NL OA PT SD SE SZ  
W: AM AU BB BG BR BY CA CN CZ FI GE HU JP KE KG KP KR KZ LK LT LV MD MG MN MW NO NZ PL RO RU SD SI SK TJ TT UA UZ VN  
AU 9512596 A 950613 (9539)  
ADT WO 9514714 A1 WO 94-US13542 941122; AU 9512596 A AU 95-12596 941122  
FDT AU 9512596 A Based on WO 9514714  
PRAI US 94-286861 940804; US 93-158001 931124  
AN 95-206899 [27] WPIDS  
AB WO 9514714 A UPAB: 950712

The following are claimed: (A) a peptide that binds to the alpha5beta1 integrin, comprising the sequence: RX1ETX2WX3(I) where X1, X2 and X3 are amino acids; (B) a peptide that binds alpha5beta1 integrin and that contains RGDGX, where X is an amino acid with a hydrophobic, aromatic side chain; (C) a peptide that binds to alphavbeta3 integrin and that contains the sequence RLD is a constrained sec. conformation; (D) a peptide that binds to the alphavbeta3 and alphavbeta5 integrins and that contains the sequence X1X2X3RGDX4X5X6, where X1, X3, X4 and X6 are capable of forming a bridge and X2 and X5 are 1-5 amino acids; (E) devices comprising the peptides of (A), (B), (C) and (D) attached to the surface of a substrate, pref. the surface of an **implantable** prosthetic; and (F) patch grafts comprising the peptides of (A), (B) (C) and (D) attached to a support matrix.

USE - The peptides are useful for isolating the complementary integrin (alpha5 or betav contg.) from a sample mixt. by contacting it under ionic conditions to allow binding of the integrin to the peptide, and then sepg. the integrin from the peptide. They can also be used for attaching cells to a substrate, by binding them to the substrate of interest and then contacting the substrate with the cell. In addn., the peptides can be used for attracting cells to the surface of an **implantable** prosthetic. The peptides also promote wound healing, when applied locally and inhibit the attachment of osteoclasts to bone. They can also be used to inhibit angiogenesis, metastasis of a tumour and migration of smooth muscle cells (all claimed). Where the treatment is localised, the peptide is administered coupled to a suitable carrier such as **hyaluronic acid**, which can be given **topically**.

When treatment is systemic, the compsn. can be administered parenterally e.g. intravenously, intramuscularly, subcutaneously, intraorbitally, intracapsularly, intraperitoneally or intracisternally.

ADVANTAGE - The peptides have high binding affinities and therefore smaller doses can be given than for other RGD-contg. peptides. They also bind their target, integrins, more selectively and specifically.

Dwg.4/13

L86 ANSWER 6 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD  
 AN 95-194399 [26] WPIDS  
 DNC C95-089973  
 TI Compsn. for preventing stenosis of tubular walls - contains hyaluronic acid and opt. vitamin C, NSAID, restenosis inhibitor, antioxidant or free radical scavenger.  
 DC B04 B05  
 IN **ASCULAI, S S; FALK, R E; TURLEY, E A**  
 PA (NORP-N) NORPHARMCO INC  
 CYC 1  
 PI CA 2106695 A 950323 (9526)\* 42 pp  
 ADT CA 2106695 A CA 93-2106695 930922  
 PRAI CA 93-2106695 930922  
 AN 95-194399 [26] WPIDS  
 AB CA 2106695 A UPAB: 950705  
 Preventing narrowing of tubular walls of an animal after they have been traumatised comprises administering hyaluronic acid (HA) and/or its salts, homologues, analogues, derivs., complexes, esters, fragments and/or subunits. Also claimed is a method further comprising the admin. of a NSAID or vitamin C, an anti-oxidant, a free radical scavenger and/or a stenosis inhibitor  
 USE - The compsn. is used to prevent narrowing (stenosis) of tubular walls of an animal after they have been traumatised, partic. arterial restenosis after balloon angioplasty in humans.  
 ADVANTAGE - When 1-2 mg/kg NSAID with 200 mg HA is administered no major toxic side effects occur such as gastrointestinal distress, neurological abnormalities or depression.  
 Dwg.0/6

L86 ANSWER 7 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD  
 AN 95-082010 [11] WPIDS  
 DNC C95-036806  
 TI Compsns for **topical** treatment of burns, wounds etc. contain e.g. **hyaluronic** acid derivs - for accelerated repair of tissue.  
 DC B04  
 IN **BENEDETTI, L; CALLEGARO, L**  
 PA (FIDI-N) FIDIA ADVANCED BIOPOLYMERS SRL  
 CYC 56  
 PI WO 9503786 A2 950209 (9511)\* EN 23 pp  
 RW: AT BE CH **DE DK** ES FR GB GR IE IT KE LU MC MW NL OA PT SD SE  
 W: AM AT AU BB BG BR BY CA CH CN CZ DE DK ES FI GB GE HU JP KE  
 KG KP KR KZ LK LT LU LV MD MG MN MW NL NO NZ PL PT RO RU SD  
 SE SI SK TJ TT UA US UZ VN  
 AU 9475341 A 950228 (9521)  
 ADT WO 9503786 A2 WO 94-EP2536 940729; AU 9475341 A AU 94-75341 940729  
 FDT AU 9475341 A Based on WO 9503786  
 PRAI IT 93-PD165 930730  
 AN 95-082010 [11] WPIDS  
 AB WO 9503786 A UPAB: 950322  
 Compsn. (I) for **topical** admin. comprises an acidic polysaccharide, a gaseous vehicle and an acceptable carrier or excipient.  
 USE - (I) are useful in the treatment of skin ulcers, sores, wounds and burns to hasten healing and repair of tissues. Additional drugs e.g. antimicrobials, antifungals, etc. may be included in (I) to increase its effectiveness. The gaseous vehicle affords even distribution of the active ingredients ensuring good contact with the treatment site and the opportunity to modulate the dosage according to the severity of the injury. (I) may interact with wound exudate forming a coating which provides additional protection against **infection**. Admin. is **topical** via single or multiple dose sprays conveying aerosol, liquid, foam or dry powder compsns.  
 Dwg.0/0

L86 ANSWER 8 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD  
 AN 95-067659 [10] WPIDS  
 DNC C95-029924



TI Compsns. for treatment of cancers - comprising an NSAID, hyaluronic acid and opt. vitamin C.  
 DC B05 D21 E19  
 IN **ASCULAI, S S; FALK, R E**  
 PA (NORP-N) NORPHARMCO INC  
 CYC 1  
 PI CA 2097892 A 941207 (9510)\* 185 pp  
 ADT CA 2097892 A CA 93-2097892 930607  
 PRAI CA 93-2097892 930607  
 AN 95-067659 [10] WPIDS  
 AB CA 2097892 A UPAB: 950314

The following are claimed: (A) methods of (i) conditioning the human immune system to resist the formation of one or more cancerous tissue types, or (ii) preventing the spread and/or metastasis of one or more cancerous tissue types in humans, comprising admin. of a compsn. comprising: (a) pharmaceutical excipients; (b) an NSAID; (c) hyaluronic acid (and/or salts, homologues, analogues, derivs, complexes, esters, fragments and/or sub-units of hyaluronic acid); and opt. (d) vitamin C. (B) sunscreen compsns. including a plurality of dosage amts. of a compsn. for admin. to humans (for purposes (i) and (ii)) above), each dosage amt. comprising (a) sunscreen agents with an acceptable SPF No.; and (b) components (b), (c) and opt. (d) (as described above).

USE - The methods may be used to treat, e.g., basal cell carcinoma, squamous cell tumours, metastatic cancer of the breast to the skin, malignancies and/or tumours in the skin, primary and metastatic melanoma in skin, genital warts, cervical cancer, psoriasis, corns and hair loss on the head of pregnant women.  
 Dwg.0/10

L86 ANSWER 9 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD  
 AN 94-341470 [42] WPIDS  
 DNC C94-155499

TI Compsn. for inhibition, control and regression of angiogenesis - comprises non-steroidal antiinflammatory agent and hyaluronic acid, useful for treating e.g. sub-retinal neovascularisation, arthritis etc..

DC B04 B05  
 IN ALAM, C; **ASCULAI, S S; FALK, R E**; HARPER, D W;  
 WILLOUGHBY, D A; ALLUM, C  
 PA (NORP-N) NORPHARMCO INC  
 CYC 54  
 PI WO 9423725 A1 941027 (9442)\* EN 46 pp

RW: AT BE CH DE DK ES FR GB GR IE IT LU MC NL OA PT SE  
 W: AT AU BB BG BR BY CA CH CN CZ DE DK ES FI GB GE HU JP KG KP  
 KR KZ LK LU LV MD MG MN MW NL NO NZ PL PT RO RU SD SE SI SK  
 TJ TT UA US UZ VN

CA 2094203 A 941017 (9503)  
 AU 9465616 A 941108 (9507)  
 ZA 9402597 A 950426 (9522) 45 pp  
 FI 9504914 A 951106 (9605)  
 NO 9504073 A 951204 (9606)

ADT WO 9423725 A1 WO 94-CA207 940415; CA 2094203 A CA 93-2094203 930416;  
 AU 9465616 A AU 94-65616 940415; ZA 9402597 A ZA 94-2597 940415; FI 9504914 A WO 94-CA207 940415, FI 95-4914 951016; NO 9504073 A WO 94-CA207 940415, NO 95-4073 951013

FDT AU 9465616 A Based on WO 9423725

PRAI CA 93-2094203 930416

AN 94-341470 [42] WPIDS

AB WO 9423725 A UPAB: 950619

Compsn. for inhibiting, controlling and/or regressing angiogenesis comprises therapeutically acceptable amts. of: (a) a non-steroidal antiinflammatory agent (NSAID); and (b) hyaluronic acid and/or its salts, homologues, analogues, derivs., complexes, esters, fragments, and sub-units of hyaluronic acid.

Pref., the hyaluronic acid is sodium hyaluronate (molecular wt. less than about 750000 daltons). The NSAID is diclofenac, diclofenac sodium, indomethacin, naproxen, (+/-)-tromethamine salt of ketorolac, ibuprofen (RTM), piroxicam (RTM), propionic acid derivs.,

acetylsalicylic acid or flunixin.

USE - The compsn. is useful for treatment of sub-retinal neovascularisation, arthritis or pannus, or tumours, and as an adjunct to cancer treatment. For a 70 kg patient, the systemic dose of NSAID, e.g. diclofenac, is 15-100 mg, or larger amts. e.g. 420 mg. For every 15 mg NSAID, about 50 mg of the hyaluronic acid is used, i.e. about 50-1050 mg. Partic. pref. is 420 mg diclofenac with 220 mg sodium hyaluronate. For topical admin., the amt. of e.g. both diclofenac sodium and sodium hyaluronate is in excess of 5-10 mg/cm2 of skin or exposed tissue. Treatment is administered daily for a number of weeks.  
Dwg.0/4

L86 ANSWER 10 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD

AN 94-310921 [39] WPIDS

DNC C94-141332

TI Compsn for topical administration - contg drug, and hyaluronic acid and/or its salts and/or homologues, analogues, derivs, complexes, ester(s), fragments or sub-units.

DC B04

IN **ASCULAI, S S; FALK, R E**

PA (NORP-N) NORPHARMCO INC

CYC 1

PI CA 2089635 A 940817 (9439)\* 117 pp

ADT CA 2089635 A CA 93-2089635 930216

PRAI CA 93-2089635 930216

AN 94-310921 [39] WPIDS

AB CA 2089635 A UPAB: 941122

A pharmaceutical compsn. comprises dosage amts. of a compsn. for topical administration to the site of pathology and/or trauma of skin and/or exposed tissue of a human patient. Each dosage amt. comprises (a) a drug and (b) hyaluronic acid (HA) and/or its salts and/or homologues, analogues, derivs., complexes, esters, fragments and/or sub-units of HA to transport the drug to the site of the pathology and/or trauma.

USE/ADVANTAGE - The compsns. can be used to treat e.g. basal cell carcinoma, precancerous actinic keratoses lesions, fungal lesions, liver spots, squamous cell tumours, metastatic cancer of the breast to the skin, primary and metastatic melanoma in the skin, malignancies and/or tumours in the skin, genital warts, cervical cancer, human papilloma virus (HPV), psoriasis, corns on the feet, hair loss on the head of pregnant women or pain (claimed). The compsns. are quickly transported in dosage amts. percutaneously at a site in need of treatment and remain at the site for a prolonged period of time. The compsns. subsequently clear through the lymphatics thereby bringing dosage amts. of the compsns. to the lymphatics for the treatment of diseases and conditions in the lymphatics. Side effects and toxicity associated with the use of the drugs is reduced.  
Dwg.0/7

L86 ANSWER 11 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD

AN 94-310920 [39] WPIDS

DNC C94-141331

TI Compsn for topical administration - contg a drug and a hyaluronic acid component for rapid delivery and prolonged activity..

DC B04

IN **ASCULAI, S S; FALK, R E; HARPER, D W; HOCHMAN,**

D; KLEIN, E S; PURSCHKE, D

PA (NORP-N) NORPHARMCO INC

CYC 1

PI CA 2089621 A 940817 (9439)\* 118 pp

ADT CA 2089621 A CA 93-2089621 930216

PRAI CA 93-2089621 930216

AN 94-310920 [39] WPIDS

AB CA 2089621 A UPAB: 941122

(A) Pharmaceutical compsns. from which effective non-toxic dosage amts. may be taken and applied to the skin and/or exposed tissue of a human are claimed. Each effective dosage amt. comprises excipients

for topical application, a drug to treat and to assist to resolve a disease and/or condition of the skin and/or exposed tissue of a human and hyaluronic acid (HA) and/or its salts and/or homologues, analogues, derivs., complexes, esters, fragments and/or sub-units of HA to transport (to facilitate or cause the transport of) the drug to a site in the skin including epidermis or exposed tissue of a disease or condition for percutaneous transport into the skin and/or exposed tissue to accumulate and remain there for a prolonged period of time and which is systemic independent acting.

USE/ADVANTAGE - The compsns. can be used for treating e.g. basal cell carcinoma, precancerous actinic keratoses lesions, fungal lesions, liver spots, squamous cell tumours, metastatic cancer of the breast to the skin, primary and metastatic melanoma in the skin, malignancies and/or tumours of the skin, genital warts, cervical cancer, human papilloma virus (HPV), psoriasis, corns on the feet or hair loss on the head of pregnant women (claimed). The compsns. are quickly transported in dosage amts. percutaneously at a site in need of treatment and remain at the site for a prolonged period of time. The compsns. subsequently clear through the lymphatics thereby bringing dosage amts. of the compsns. to the lymphatics for the treatment of disease and conditions in the lymphatics. Side effects and toxicity associated with the use of the drugs is reduced.  
Dwg.0/7

L86 ANSWER 12 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD  
AN 94-135209 [16] WPIDS  
DNC C94-062504  
TI Prevention of the narrowing of tubular walls after trauma - by administering hyaluronic acid, esp. suitable for preventing arterial restenosis after e.g. balloon angioplasty..  
DC B04  
IN **ASCULAI, S S; FALK, R E**; TURLEY, E A; SCULAI, S  
PA (NORP-N) NORPHARMCO INC  
CYC 46  
PI WO 9407505 A1 940414 (9416)\* EN 61 pp  
RW: AT BE CH DE DK ES FR GB GR IE IT LU MC NL OA PT SE  
W: AT AU BB BG BR BY CA CH CZ DE DK ES FI GB HU JP KP KR KZ LK  
LU MG MN MW NL NO NZ PL PT RO RU SD SE SK UA US VN  
CA 2079205 A 940326 (9423)  
ZA 9307068 A 940629 (9428) 42 pp  
AU 9348126 A 940426 (9432)  
NO 9501122 A 950323 (9524)  
EP 661981 A1 950712 (9532) EN  
R: AT BE CH DE DK ES FR GB GR IE IT LI LU MC NL PT SE  
ADT WO 9407505 A1 WO 93-CA388 930922; CA 2079205 A CA 92-2079205 920925;  
ZA 9307068 A ZA 93-7068 930924; AU 9348126 A AU 93-48126 930922; NO 9501122 A WO 93-CA388 930922, NO 95-1122 950323; EP 661981 A1 EP 93-920624 930922, WO 93-CA388 930922  
FDT AU 9348126 A Based on WO 9407505; EP 661981 A1 Based on WO 9407505  
PRAI CA 92-2079205 920925  
AN 94-135209 [16] WPIDS  
AB WO 9407505 A UPAB: 940608

The narrowing of tubular walls of an animal after the tubular walls have been traumatised can be prevented by administering a therapeutically effective, non-toxic amt. of hyaluronic acid and/or its salts and/or homologous, analogues, derivs. complexes, esters, in fragments or subunits.

USE - The hyaluronic acid can prevent narrowing of tubular walls after they have been traumatised and is esp. useful in preventing arterial restenosis after e.g. balloon angioplasty when endothelial cell proliferation occurs on the inner arterial wall caused by irritation to the cells by the balloon angioplasty. The hyaluronic acid is safe and essentially non-toxic. The hyaluronic acid is administered i.v. in amts. of 10-3000 mg for a 70kg person, prior to, during and/or after injury.

Dwg.0/0

L86 ANSWER 13 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD

AN 93-382262 [48] WPIDS  
 DNN N93-295528 DNC C93-169391  
 TI Artificial skin, protecting surface of wound and preventing pain -  
 comprises wound contact layer including microcapsule comprising heat  
 modified collagen and muco-polysaccharide and layer adjusting water  
 vapour permeation.  
 DC B04 B07 D22 P32 P34  
 PA (TERU) TERUMO CORP  
 CYC 1  
 PI JP 05285210 A 931102 (9348)\* 6 pp  
 ADT JP 05285210 A JP 92-94328 920414  
 PRAI JP 92-94328 920414  
 AN 93-382262 [48] WPIDS  
 AB JP05285210 A UPAB: 940120

Skin comprises a wound contact layer comprising collagen matrix  
 contg. a microcapsule including a substance promoting the prodn. of  
 collagen and a layer adjusting the permeation of water vapour. The  
 microcapsule has a coacervate structure comprising heat-modified  
 collagen and mucopolysaccharide(s).

The collagen matrix pref. comprises fibrous collagen and a  
 modified collagen of a helix content of 0-80 (0-50)%. The substance  
 promoting the prodn. of collagen is (A) ascorbic acid and/or its  
 phosphoric ester(s) or (B) chitin and/or its deriv(s).. The  
 mucopolysaccharides include chondroitin, **hyaluronic acid**  
 and heparin.

USE/ADVANTAGE - The skin protects the surface of wounds softly  
 and prevents pain and **infection**. It allows early cell  
 invasion, promoting invasion and growth of fibroblasts. The upper  
 adjusting layer may be peeled off after a specific period to  
**implant self-graft skin.**  
 Dwg.0/0

L86 ANSWER 14 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD

AN 93-336584 [42] WPIDS

DNC C93-148866

TI Non-fibrotic growth factor and opt. anti-fibrotic agent compsn. -  
 utilised for stimulating wound healing without fibrosis, also for  
 treating fibrotic disease.

DC B04 D16

IN FERGUSON, M W J; SHAH, M; SHAH, H

PA (UYMA-N) UNIV VICTORIA MANCHESTER

CYC 42

PI WO 9319769 A1 931014 (9342)\* EN 26 pp

RW: AT BE CH DE DK ES FR GB GR IE IT LU MC NL OA PT SE

W: AT AU BB BG BR CA CH CZ DE DK ES FI GB HU JP KP KR LK LU MG

MN MW NL NO NZ PL PT RO RU SD SE SK UA US

AU 9337623 A 931108 (9408)

NO 9403466 A 940916 (9443)

EP 646012 A1 950405 (9518) EN

R: AT BE CH DE DK ES FR GB GR IE IT LI LU MC NL PT SE

CZ 9402366 A3 950412 (9524)

SK 9401168 A3 950208 (9525)

JP 07505378 W 950615 (9532)

HU 68905 T 950828 (9540)

ADT WO 9319769 A1 WO 93-GB586 930322; AU 9337623 A AU 93-37623 930322;  
 NO 9403466 A WO 93-GB586 930322, NO 94-3466 940916; EP 646012 A1 EP  
 93-906723 930322, WO 93-GB586 930322; CZ 9402366 A3 CZ 94-2366  
 930322; SK 9401168 A3 SK 94-1168 940928, WO 93-GB586 ; JP  
 07505378 W JP 93-517193 930322, WO 93-GB586 930322; HU 68905 T WO  
 93-GB586 930322, HU 94-2771 930322

FDT AU 9337623 A Based on WO 9319769; EP 646012 A1 Based on WO 9319769;  
 JP 07505378 W Based on WO 9319769; HU 68905 T Based on WO 9319769

PRAI GB 92-6861 920328

AN 93-336584 [42] WPIDS

AB WO 9319769 A UPAB: 931202

Healing compsn. (A) contains at least one non-fibrotic growth factor  
 (I) and a pharmaceutically acceptable carrier.

Esp. (I) is transforming growth factor (TGF) beta 3 (Ia) or  
 fibroblast growth factor (Ib) and the compsn. may include an

anti-fibrotic agent (II). (I) and (II) can be present in active or inactive form (e.g. in a capsule which can be degraded by UV light, ultrasound, in vivo enzymes or heat).

USE/ADVANTAGE - (A) is used to facilitate repair and healing of wounds without excessive fibrosis and also to treat fibrotic conditions (e.g. liver cirrhosis, glomerulonephritis, pulmonary fibrosis, ulcers, etc.). (I) is formulated with a neutral sterile cream, gel, aerosol or powder for **topical** application; as a patch or dressing; as a sterile soln. for irrigation, injection or inhalation, or as a tablet or capsule. The carrier may also be a biopolymer (e.g. collagen or **hyaluronic acid**) for use as an **implant** or controlled release system.

Dwg.0/0

L86 ANSWER 15 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD

AN 93-288135 [36] WPIDS

DNC C93-128581

TI Topical formulations contg. hyaluronic acid (deriv.) - used for promoting transport of drug, esp. antiinflammatory or anticancer agent, into skin, exposed tissue or lymphatic system.

DC B04 B07

IN ASCULAI, S S; FALK, R E

PA (NORP-N) NORPHARMCO INC; (HYAL-N) HYAL PHARM CORP

CYC 43

PI WO 9316733 A1 930902 (9336)\* EN 106 pp

RW: AT BE CH DE DK ES FR GB GR IE IT LU MC NL OA PT SE

W: AT AU BB BG BR CA CH CZ DE DK ES FI GB HU JP KP KR LK LU MG

MN MW NL NO NZ PL PT RO RU SD SE SK UA US

CA 2061566 A 930821 (9345)

ZA 9301174 A 931124 (9402) 120 pp

AU 9334889 A 930913 (9403)

SK 9300110 A3 930909 (9419)

EP 626864 A1 941207 (9502) EN

R: AT BE CH DE DK ES FR GB GR IE IT LI LU MC NL PT SE

JP 07507054 W 950803 (9539) 37 pp

ADT WO 9316733 A1 WO 93-CA62 930216; CA 2061566 A CA 92-2061566 920220;

ZA 9301174 A ZA 93-1174 930219; AU 9334889 A AU 93-34889 930216; SK

9300110 A3 SK 93-110 930222; EP 626864 A1 EP 93-903755 930216, WO

93-CA62 930216; JP 07507054 W JP 93-514408 930216, WO 93-CA62 930216

FDT AU 9334889 A Based on WO 9316733; EP 626864 A1 Based on WO 9316733;

JP 07507054 W Based on WO 9316733

PRAI CA 92-2061566 920220

AN 93-288135 [36] WPIDS

AB WO 9316733 A UPAB: 950804

Pharmaceutical compsn. for topical applicatin to human skin or exposed tissue contains sufficient drug (I) to treat a condition of the skin and/or exposed tissue and sufficient hyaluronic acid (HA) (and/or its salts, homologues, analogues, derivs., complexes, esters and/or subunits) to facilitate transport of (I) into the desired site for treatment.

Specifically (I) remains and accumulates in the desired region for a proloned period. It may be discharged via the lymphatic system (to treat conditions of the lumphatics).

USE/ADVANTAGE - The condition treated is specifically basal cell carcinoma, precancerous (often recurrent) actinic keratoses lesions, fungal lesions, liver spots, squamous cell tumours, metastatic cancer of the breast to the skin, prim. and metastatic melanoma in the skin, malignancy and/or tumour of the skin, genital warts (condyloma acumirata), cerical cancer, HPV (human papilloma virus, e.g. of the cervic), psoriasis (plaque-type or nail bed, corns on the feet of hair loss in pregnant women. Dose of HA or deriv. is at least 5-10 mg (cm2. The compsn. is rubbed into the desired region once or a few times daily for a period of weeks. The treatment may involve blocking prostaglanolin, synthesis to enable macrophages and NK cells to resolve the condition. Alternatively, (I) relieves pain by transport into the epidermis adjacent to the paccian nerve bundle compsns. are systemic independent, i.e. (I) does not enter the blood stream. (I) are selectively and rapidly targetted to the desired site of action epidermis), providing

improved therapeutic effect and reduced toxicity and side-effects.

Dwg.0/7

Dwg.0/7

L86 ANSWER 16 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD

AN 93-288134 [36] WPIDS

DNC C93-128580

TI Topical formations contg. hyaluaronic acid (deriv.) - used for promoting transport of drug, esp. antiinflammatory or anticancer agent, into skin, exposed tissue or lymphatic system.

DC B04 B07

IN ASCULAI, S S; FALK, R E; HARPER, D W; HOCHMAN, D; KLEIN, E S; PURSCHKE, D

PA (NORP-N) NORPHARMCO INC; (HYAL-N) HYAL PHARM CORP

CYC 44

PI WO 9316732 A1 930902 (9336)\* EN 107 pp

RW: AT BE CH DE DK ES FR GB GR IE IT LU MC NL OA PT SE

W: AT AU BB BG BR CA CH CZ DE DK ES FI GB HU JP KP KR LK LU MG

MN MW NL NO NZ PL PT RO RU SD SE SK UA US

CA 2061703 A 930821 (9345)

AU 9334888 A 930913 (9403)

SK 9300111 A3 930909 (9419)

EP 626863 A1 941207 (9502) EN

R: AT BE CH DE DK ES FR GB GR IE IT LI LU MC NL PT SE

FI 9403789 A 941003 (9502)

NO 9403044 A 941019 (9502)

ZA 9301173 A 950222 (9514) 154 pp

CN 1084064 A 940323 (9525)

JP 07506812 W 950727 (9538) 45 pp

ADT WO 9316732 A1 WO 93-CA61 930216; CA 2061703 A CA 92-2061703 920220;

AU 9334888 A AU 93-34888 930216; SK 9300111 A3 SK 93-111 930505; EP

626863 A1 EP 93-903754 930216, WO 93-CA61 930216; FI 9403789 A WO

93-CA61 930216, FI 94-3789 940817; NO 9403044 A WO 93-CA61 930216,

NO 94-3044 940817; ZA 9301173 A ZA 93-1173 930219; CN 1084064 A CN

93-103488 930220; JP 07506812 W JP 93-514407 930216, WO 93-CA61

930216

FDT AU 9334888 A Based on WO 9316732; EP 626863 A1 Based on WO 9316732;

JP 07506812 W Based on WO 9316732

PRAI CA 92-2061703 920220

AN 93-288134 [36] WPIDS

AB WO 9316732 A UPAB: 950804

Pharmaceutical compsn. for application to human skin and/or exposed tissue contains topical excipients, sufficient drug (I) to treat a condition of the skin and/or exposed tissue and sufficient hyaluronic acid (HA) (and/or its salts, homologues, analogues, derivs., complexes, esters, fragments and/or subunits) to facilitate transport of (I) to a suitable site in the skin (including epidermis) and/or exposed tissue for percutaneous transport into the desired treatment. (I) accumulator in the required site, remains for a prolonged period and is system indepent acting.

The HA component is HA or its salt, having mol. wt. less than 750000 daltons. (I) is pref. a non-steroidal antiinflammatory drug (NSAID) (esp. dichlofenac, indomethacin, naproxen, (+)-tromethamine salt of Potorolac, ibuprofen, piroxicam, propionic acid deriv., acetylsalicyclio acid or flunixin) or an anticancer drug (esp. novantrone or 5-fluorouracil).

USE/ADVANTAGE - The condition treated is specifically basal cell carcinoma, precancerous (often recurrent) actinic keratoses lesions, fungal lesions, liver spots, squamous cell tumours, metastatic cancer of the breast to the skin, prim. and metastatic melanoma in the skin, malignacny and/or tumour of the skin, genital worts, cervical cancer, HPV (human papilloma virus, e.g. of the cervic), psoriasis (plaque-type or nail bed), corns on the feet or hair loss in pregnant women. Dose of HA or deriv. is more than 5 mg per sq. cm. (I) are selectively and rapidly targetted at the desired site of action, providing improved therapeutic effect and reduced toxicity and adverse effects typically (I) remain at the desired site for more than 12-24 hrs..

Dwg.0/7

Dwg.0/7

L86 ANSWER 17 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD  
 AN 93-266414 [34] WPIDS  
 DNC C93-118709  
 TI Hyaluronic acid (salts) and derivs. - are used in pharmaceutical  
 compsn. for treating ischaemia, damage in tissue.  
 DC B04  
 IN ASCULAI, S S; FALK, R E; KLEIN, E S  
 PA (NORP-N) NORPHARMCO INC; (HYAL-N) HYAL PHARM CORP  
 CYC 18  
 PI EP 557118 A1 930825 (9334)\* EN 19 pp  
 R: AT BE CH DE DK ES FR GB GR IE IT LI LU MC NL PT SE  
 CA 2061567 A 930821 (9345)  
 ADT EP 557118 A1 EP 93-301230 930219; CA 2061567 A CA 92-2061567 920220  
 PRAI CA 92-2061567 920220  
 AN 93-266414 [34] WPIDS  
 AB EP 557118 A UPAB: 950602  
 A pharmaceutical compsn. comprises hyaluronic acid (HA) and/or salts  
 of HA and/or homologues, analogues, derivs., complexes, esters,  
 fragments and units of HA in association with a diluent or carrier.  
 USE - Alanine Aminotransferase prodn. in damaged tissue is  
 reduced by administration of the compsn., which has use in  
 preventing or repairing ischaemia reperfusion damage in tissue,  
 partic. internal organs, e.g., the liver, kidneys and heart. Thus it  
 may be used to treat ischaemia damage in tissue during  
 transplantation. Pref. HA and its salts are used in amts. of 300mg  
 to 7g per day for a 70kg human.  
 Dwg.0/4  
 Dwg.0/4

L86 ANSWER 18 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD  
 AN 93-134088 [16] WPIDS  
 DNN N93-102273 DNC C93-059798  
 TI Flexible wound dressing prepn. for keeping wound in moist condition  
 - by mixing dry hydrocolloid polymer e.g. guar or xanthan gum with  
 water in a sealed package to form dispersion which is then  
 solidified.  
 DC A96 B07 D22 P32  
 IN ROLF, D  
 PA (ROLF-I) ROLF D; (LECT-N) LECTEC CORP  
 CYC 25  
 PI WO 9306802 A1 930415 (9316)\* EN 41 pp  
 RW: AT BE CH DE DK ES FR GB GR IE IT LU MC NL SE  
 W: AU BR CA FI JP KR NO RU  
 ZA 9207749 A 930630 (9332) 43 pp  
 AU 9227857 A 930503 (9334)  
 NO 9401284 A 940608 (9429)  
 FI 9401627 A 940608 (9431)  
 EP 625894 A1 941130 (9501) EN  
 R: AT BE CH DE DK FR GB IE IT LI LU NL SE  
 JP 07500035 W 950105 (9511)  
 AU 663737 B 951019 (9549)  
 ADT WO 9306802 A1 WO 92-US8403 921002; ZA 9207749 A ZA 92-7749 921008;  
 AU 9227857 A AU 92-27857 921002; NO 9401284 A WO 92-US8403 921002,  
 NO 94-1284 940408; FI 9401627 A WO 92-US8403 921002, FI 94-1627  
 940408; EP 625894 A1 EP 92-921920 921002, WO 92-US8403 921002; JP  
 07500035 W WO 92-US8403 921002, JP 93-507081 921002; AU 663737 B AU  
 92-27857 921002  
 FDT AU 9227857 A Based on WO 9306802; EP 625894 A1 Based on WO 9306802;  
 JP 07500035 W Based on WO 9306802; AU 663737 B Previous Publ. AU  
 9227857, Based on WO 9306802  
 PRAI US 91-774064 911009; US 92-913151 920714; US 92-914751 920715  
 AN 93-134088 [16] WPIDS  
 AB WO 9306802 A UPAB: 931115  
 The wound dressing is prepd. as follows: (a) a dry, particulate  
 H2O-soluble or -swellable natural or synth. hydrocolloid polymer (I)  
 is sealed in a package in a sterile state and kept in a dry  
 condition; (b) (I) is hydrated before use by mixing it with H2O

within the sealed package to afford a fluid dispersion which can be poured from the package or spread onto a surface to conform with it; and (c) the dispersion is allowed to solidify to furnish a solid but flexible hydrated gel dressing.

Pref. (I) are guar gum or a deriv. (pref. cationic guar, hydroxy propyl guar, and anionic guar), galactomannan, glucomannan, xanthan gum, locust bean gum, algin or mixts. pref. a crosslinking agent (II) is also present to enhance gelling of (I) (esp. H3BO3, borax, an organic titanate, galactose, mannose, lactose, a galactose- or mannose-contg. oligosaccharide, or a source of H2O-soluble Ca, Mg or Al cations or mixts.), as well as a medicament (esp. medication, **disinfectant**, wound healing enhancer, vitamin, blood coagulant, antibiotic, or O2 source or mixts., esp. coagulant, alum, witch hazel, neomycin sulphate, bacitracin, **hyaluronic acid** (or fragment for promoting pathogenic wound healing), an analgesic, morphine, fentanyl, lidocaine, procaine, and epinephrine).

In the package, (I) and the H2O are pref. contained in 2 separate compartments of the package, with a rupturable membrane between them; (I) is pref. sterilised by a gas (entry through a porous polytetrafluoroethylene portion to allow entry), and the H2O is pref. sterilised by another means, e.g. irradiation. pref. dressings comprise (by wt.) 3-15% (I) (esp. 8-15% guar gum), and 0.1-5% (II) (esp. 0.1-1% H3BO3).

USE/ADVANTAGE - The produced flexible hydrated gel dressing keeps wounds in a moist condition, and absorbs exudate from them and cushions them. The dressing is quickly prepd. and applied from shelf-stable components which need not refrigeration. The dressing is supple, elastic, pliable, and soft, keeps its shape thorough a wide range of temps., and can be removed from the wound bed as a solid plug.

1/10

Dwg.1/10

ABEQ ZA 9207749 A UPAB: 931118

A water-based natural or synthetic hydrocolloidal polymeric gel comprises a gel-forming hydrocolloid polymer in dry particulate form and a source of water or an aq. soln. The liq. and dry solid components are initially separate and are typically contained in separate compartments of a sealed package but are mixed together within the package, e.g., a flexible pouch, just before use. The hydrocolloid does not become fully hydrated immediately. The liq. component gives the mixt. a fluid consistency when mixed with the hydrocolloid. At this stage, the admixture is sufficiently fluid in consistency to allow it to be poured or spread into the wound. Following application to the wound, the hydrated hydrocolloidal dispersion begins to solidify to form a self-supporting, solid but flexible dressing structure consisting primarily of water and the hydrocolloid. A gelling or crosslinking agent can also be used if desired with some of the hydrocolloids. Opt. the dressing can also contain a biologically active agent, e.g., a medicament.

USE/ADVANTAGE - Used for dressing wounds and **implantation** beneath the skin of a patient. The resulting dressing becomes moulded to the shape of the wound and contains a large quantity of moisture that will maintain the wound in a moist condition.

L86 ANSWER 19 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD

AN 92-315907 [38] WPIDS

DNC C92-140310

TI Liposome(s) providing sustained and targetted drug delivery - are surface modified by attaching cpd. which provides specific adhesion to target site, for treating burns, wounds, tumours, etc..

DC B05 B07

IN MARGALIT, R

PA (BAXT) BAXTER INT INC

CYC 17

PI WO 9214445 A1 920903 (9238)\* EN 20 pp

RW: AT BE CH DE DK ES FR GB GR IT LU NL SE

W: AU CA JP



AU 9213677 A 920915 (9251)  
 EP 525167 A1 930203 (9305) EN 20 pp  
 R: AT BE CH DE DK ES FR GB IT LI LU NL SE  
 JP 05506253 W 930916 (9342) 6 pp  
 EP 525167 B1 950913 (9541) EN 8 pp  
 R: AT BE CH DE DK ES FR GB IT LI LU NL SE  
 DE 69113036 E 951019 (9547)

ADT WO 9214445 A1 WO 91-US8111 911030; AU 9213677 A WO 91-US8111 911030,  
 AU 92-13677 911030; EP 525167 A1 WO 91-US8111 911030, EP 92-906507  
 911030; JP 05506253 W WO 91-US8111 911030, JP 92-505852 911030; EP  
 525167 B1 WO 91-US8111 911030, EP 92-906507 911030; DE 69113036 E DE  
 91-613036 911030, WO 91-US8111 911030, EP 92-906507 911030

FDT AU 9213677 A Based on WO 9214445; EP 525167 A1 Based on WO 9214445;  
 JP 05506253 W Based on WO 9214445; EP 525167 B1 Based on WO 9214445;  
 DE 69113036 E Based on EP 525167, Based on WO 9214445

PRAI US 91-655879 910214  
 AN 92-315907 [38] WPIDS  
 AB WO 9214445 A UPAB: 931113

Microscopic delivery system for sustained release of a substance (I) comprises a liposome component, (I) encapsulated by the liposome, and a recognition substance (II) bonded to the liposome surface.

The liposome component is pref. multilamellar or large unilamellar vesicle, or a microemulsified liposome, esp. it includes phosphatidyl ethanolamine. (II) is gelatin, collagen, **hyaluronic** acid or epidermal growth factor and is covalently bonded to the liposome, esp. through a crosslinking agent.

USE/ADVANTAGE - (II) provides target specifically for and retention on, partic. cellular to sites where (I) is required to be released. The new system is superior to conventional liposomes (which cannot be retained at the target site) and attachment of (II) has no significant effect on drug delivery. The new system is delivered **topically**, e.g. in treatment of burns, wounds, **infections**, tumours etc.

Dwg.0/0

ABEQ JP05506253 W UPAB: 931202

Microscopic delivery system for sustained release of a substance (I) comprises a liposome component, (I) encapsulated by the liposome, and a recognition substance (II) bonded to the liposome surface.

The liposome component is pref. multilamellar or large unilamellar vesicle, or a microemulsified liposome, esp. it includes phosphatidyl ethanolamine. (II) is gelatin, collagen, **hyaluronic** acid or epidermal growth factor and is covalently bonded to the liposome, esp. through a crosslinking agent.

USE/ADVANTAGE - (II) provides target specifically for and retention on, partic. cellular to sites where (I) is required to be released. The new system is superior to conventional liposomes (which cannot be retained at the target site) and attachment of (II) has no significant effect on drug delivery. The new system is delivered **topically**, e.g. in treatment of burns, wounds, **infections**, tumours etc.

ABEQ EP 525167 B UPAB: 951019

A microscopic delivery system for the sustained release of a substance comprising a liposome component, a substance encapsulated by the liposome component and a target-recognizing component covalently bonded to the liposomal surface the liposome component having a permeability that allows sustained substance release, characterised in that the target-recognizing component is selected from gelatin, collagen and **hyaluronic** acid.

Dwg.0/0

L86 ANSWER 20 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD  
 AN 92-124904 [16] WPIDS  
 DNC C92-058251  
 TI Compsn. for treating lesions, sores, ulcerations and burns - comprises **hyaluronic** acid sodium salt, hexetidine, sulphadiazine silver or zinc salts.  
 DC B04 B05  
 IN DONATPEDE, E  
 PA (ALTE-N) ALTERGON SA

CYC 7

PI EP 480189 A 920415 (9216)\* EN 7 pp

R: BE CH DE FR GB IT LI

IT 1243435 B 940610 (9441)

ADT EP 480189 A EP 91-115360 910911; IT 1243435 B IT 90-21662 901005

PRAI IT 90-21662 901005

AN 92-124904 [16] WPIDS

AB EP 480189 A UPAB: 931006

New **topical** compsn. comprises **hyaluronic** acid Nasalt and **disinfectant** chosen from gp. consisting of cresol derivs., hexetidine, sulphadiazine Ag and its Zn salt is new.

Pref. the cresol derivs. are chloroxylenol and dichloroxylenol. The excipients and diluents used include hydroxypropylmethyl-cellulose, sorbitol, glycerin, polyoxyethylenated, glycolized, glycerides, polyethyleneglycol stearate, stearic acid, oleic acid decyl ester, caprylic and caproic acid ester, ethoxylated glycerides of palmitic and lauric acids, polymerised polyvinyl alcohol, self-emulsifying wax and non-denatured collagen. The compsn. is prepd. as oil-in-water, water-in-oil emulsions, hydrogels, pastes, ointments, lotions and powders.

USE/ADVANTAGE - Admin. of exogenous **hyaluronic** acid determines an antiphlogistic and stimulating action on the granulation tissue, which accelerates cicatrization and re-epithelialization of lesions, useful in the treatment of sores, ulcerations and burns. (0/0)  
0/0

L86 ANSWER 21 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD

AN 91-275565 [38] WPIDS

DNC C91-119397

TI DNA encoding human ciliary neuronotrophic factor - is useful for treating nervous disorders such as acute or chronic pathological conditions e.g. cerebrovascular, **infective**, etc..

DC B04 D16

IN CALLEGARO, L; NEGRO, A; VALLE, F D; DELLAVALLE, F; DELLA-VALLE, F

PA (FIDI-N) FIDIA SPA

CYC 19

PI EP 446931 A 910918 (9138)\*

R: AT BE CH DE ES FR GB GR IT LI LU NL SE

AU 9172918 A 910919 (9145)

NO 9101025 A 910916 (9146)

CA 2038208 A 910915 (9149)

CN 1057295 A 911225 (9237)

JP 04218374 A 920807 (9238)

HU 62033 T 930329 (9316)

IT 1239272 B 931001 (9410)

IT 1243286 B 940526 (9441)

17 pp

ADT EP 446931 A EP 91-103969 910314; CN 1057295 A CN 91-102344 910314;

JP 04218374 A JP 91-74704 910314; HU 62033 T HU 91-840 910314; IT

1239272 B IT 90-41557 900314; IT 1243286 B IT 90-41655 900717

PRAI IT 90-41557 900314; IT 90-41655 900717

AN 91-275565 [38] WPIDS

AB EP 446931 A UPAB: 930928

A DNA isolate comprising a DNA sequence encoding human ciliary neuronotrophic factor (I) is claimed. Also new is a recombinant expression vector containing the DNA sequence, a microorganism transformed with a vector, a cell culture transformed with the vector and pure (I). The DNA sequence and the amino acid sequence of (I) are given in the specification. The microorganism is preferably E coli, and the cell line is a non-human mammalian cell line, preferably a Chinese hamster ovary cell line.

USE/ADVANTAGE - (I) is used to treat nervous disorders i.e. to maintain, prevent loss of an to treat the loss of nervous function due to acute or chronic pathological conditions, including the treatment of acute conditions e.g. cerebrovascular, **infective**, inflammatory, compressive and metabolic deficiencies, and chronic or neurodegenerative conditions. (I) is also used to treat neuropathological conditions caused by ageing of the nervous system or diseases affecting the immune system. (I) is

in the form of a composition that may also contain a natural ganglioside or its derivative, semisynthetic analogue or salt, and a natural polysaccharide, its derivative or semisynthetic analogue e.g. **hyaluronic acid**.

Administration is by subcutaneous, intramuscular or intracerebral injection, at a dosage of 0.05-5mg/kg/day. Administration can also be oral, **topical**, rectal, parenteral, local or by inhalant. @ (31pp DWg.No.0/11)DV

L86 ANSWER 22 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD

AN 91-261303 [36] WPIDS

DNC C91-113409

TI **Topical** compositions based on high-mol. weight sodium hyaluronate - used for treating inflammations of the oral cavity and for oral cavity hygiene and cosmetic treatment.

DC B04 D21

IN DISCHIENA, M G; DI, SCHIENA M G

PA (RICE-N) RICERCHÉ DI SCHIENA DI MICHELE DI SCHIEN; (RICE-N)

RICERFARMA SRL; (RICE-N) RICERCHÉ DI SCHIENA DEL MICHELE

CYC 7

PI EP 444492 A 910904 (9136)\*

R: DE ES FR GB GR IT

PT 98714 A 930226 (9312)#

IT 1240316 B 931207 (9415)

ADT EP 444492 A EP 91-102240 910218; PT 98714 A PT 91-98714 910819; IT 1240316 B IT 90-19438 900221

PRAI IT 90-19438 900221

AN 91-261303 [36] WPIDS

AB EP 444492 A UPAB: 940421

The use of **hyaluronic acid** (A) in the form of its Na salt is new. (A) has an average Mr of 800,000-4,000,000 and is used **topically** for the therapy and prophylaxis of inflammatory **infections** of the oral cavity, and for hygiene and cosmetic treatment of the oral cavity. (A) preferably has an average Mr of 1-2,000,000.

USE/ADVANTAGE - Compositions of (A) are also used to treat gingivitis, stomatitis and irritation due to mechanical causes e.g. fixed or mobile prostheses or surgical operations. General amounts in compositions are 0.005-10% by weight, with therapeutic compositions containing 0.2-10 (0.2-1)% and for prophylactic, cosmetic and hygienic treatment 0.005-0.1 (0.01)%. The compositions are in the form of gingival pastes (e.g. for the dentition stage in children), toothpastes, mouthwashes and adhesive pastes and powders.

A composition of 54g Na carboxymethylcellulose dispersed in 774g H<sub>2</sub>O (containing 0.13% p-oxybenzoate and 0.007% propyl p-oxybenzoate as preservatives), with 240g 1% Na hyaluronate (Mr 1,500,000) and 120g 70% sorbitol is prepared. 6 g peppermint is added as flavouring. This paste containing 0.2% (A) was tested on 10 patients having various degrees of parodontal pathology. By the second day, patients suffering from marginal gingivitis showed a reduction in symptomatology with complete recovery in 1 week. Slower recovery was seen in patients having undergone parodontal surgery, but a clear improvement was seen around the mucosa at the wound level, e.g. it was trophic and pink-coloured. @ (8pp Dwg.No.0/0)@

A composition of 54g Na carboxymethylcellulose dispersed in 774g H<sub>2</sub>O (containing 0.13% p-oxybenzoate and 0.007% propyl p-oxybenzoate as preservatives), with 240g 1% Na hyaluronate (Mr 1,500,000) and 120g 70% sorbitol is prepared. 6 g peppermint is added as flavouring. This paste containing 0.2% (A) was tested on 10 patients having various degrees of parodontal pathology. By the second day, patients suffering from marginal gingivitis showed a reduction in symptomatology with complete recovery in 1 week. Slower recovery was seen in patients having undergone parodontal surgery, but a clear improvement was seen around the mucosa at the wound level, e.g. it was trophic and pink-coloured.

0/0

L86 ANSWER 23 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD

AN 91-117336 [16] WPIDS

DNC C91-050471  
 TI Combinations of drug and **hyaluronic** acid - to improve tissue and cell penetration.  
 DC B05 B07 C03 D21  
 IN **ASCULAI, S S; FALK, R E**  
 PA (NORP-N) NORPHARMCO INC; (HYAL-N) HYAL PHARM CORP  
 CYC 35  
 PI WO 9104058 A 910404 (9116)\*  
 RW: AT BE CH DE DK ES FR GB IT LU NL OA SE  
 W: AT AU BB BG BR CA CH DE DK ES FI GB HU JP KP KR LK LU MC MG  
 MW NL NO RO SD SE SU US  
 AU 9064330 A 910418 (9129)  
 FI 9102470 A 910521 (9133)  
 EP 445255 A 910911 (9137)  
 R: AT BE CH DE ES FR GB IT LI LU NL SE  
 ZA 9007564 A 910828 (9139)  
 NO 9101952 A 910705 (9140)  
 BR 9006924 A 911210 (9203)  
 CN 1051503 A 910522 (9207)  
 JP 04504579 W 920813 (9239) 39 pp  
 HU 64699 T 940228 (9412)  
 AU 9352274 A 940303 (9414)  
 WO 9104058 A3 910919 (9508)  
 EP 656213 A1 950607 (9527) EN  
 R: AT BE CH DE DK ES FR GB IT LI LU NL SE  
 EP 445255 B1 951206 (9602) EN 84 pp  
 R: AT BE CH DE DK ES FR GB IT LI LU NL SE  
 ADT EP 445255 A EP 90-914108 900918; ZA 9007564 A ZA 90-7564 900921; JP 04504579 W JP 90-513204 900918; WO 90-CA306 900918; HU 64699 T HU 90-7339 900918; WO 90-CA306 900918; AU 9352274 A AU 93-52274 931209, Div ex AU 90-64330 ; WO 9104058 A3 WO 90-CA306 900918; EP 656213 A1 EP 95-100186 900918; EP 445255 B1 EP 90-914108 900918, WO 90-CA306 900918  
 FDT JP 04504579 W Based on WO 9104058; HU 64699 T Based on WO 9104058; EP 445255 B1 Based on WO 9104058  
 PRAI CA 89-612307 890921  
 AN 91-117336 [16] WPIDS  
 AB WO 9104058 A UPAB: 950602  
 New drug combinations or formulations comprise a drug and a **hyaluronic** acid cpd. (I) selected from **hyaluronic** acid and its salts, homologues, analogues, derivs., complexes, esters, fragments and subunits.  
 USE - Indications include diabetes, hormone replacement therapy, fertility control, AIDS, cancer, hair loss, herpes **infections**, renal failure, cardiac insufficiency, hypertension, oedema, microbial **infections**, acne, **transplant** rejection, inflammations, elimination of tumour breakdown material, blood detoxification, respiratory disorders, vascular ischaemia, brain tumours, mononucleosis, pain, side effects of nonsteroidal antiinflammatory agents, and tissue perfusion.  
 @ (116pp Dwg.No.0/1)  
 ABEQ EP 445255 B UPAB: 960115  
 A pharmaceutical composition comprising: (1) a medicinal and/or therapeutic agent in a therapeutically effective amount to treat a disease or condition in humans; and (2) **hyaluronic** acid and/or salts thereof and/or homologues, analogues, derivatives, complexes, esters, fragments and subunits of **hyaluronic** acid, characterised in that said composition (a) is in a dosage form which is suitable for administration in humans; and (b) is in a form in which (i) component (1) is in an effective dosage amount to treat said disease or condition by penetration at the site to be treated; and (ii) component (2) is immediately available to transport component (1) at the site to be treated, and which component (2) is in an effective non-toxic amount to facilitate the transport of component (1) upon administration, through the tissue (including scar tissue) at the site to be treated and through the cell membranes of the individual cells to be treated, wherein said amount of component (2) is sufficient to provide a dosage greater than 10 mg/70 kg person.

Dwg.0/1

L86 ANSWER 24 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD  
 AN 90-348243 [46] WPIDS  
 DNC C90-151129  
 TI Treating and/or preventing alopecia - by admin. of sulphated mono,  
 di or oligo saccharide or their derivs., salts or complexes.  
 DC B03 D21 E13  
 IN BAR-SHALOM, D; BUKH, N; BARSHALOM, D  
 PA (BUKH-N) BUKH MEDITEC AS; (BUKH-N) BUKH MEDITEC A/S; (BMRE-N) BM RES  
 AS  
 CYC 35  
 PI WO 9012561 A 901101 (9046)\*

RW: AT BE CH DE DK ES FR GB IT LU NL OA SE

W: AU BB BG BR CA FI HU JP KP KR LK MC MG MW NO RO SD SU US

AU 9055278 A 901116 (9107)

ZA 9008391 A 910925 (9144)#

FI 9104879 A 911016 (9205)

EP 469010 A 920205 (9206)

R: AT BE CH DE ES FR GB IT LI LU NL SE

NO 9104095 A 911219 (9212)

BR 9007312 A 920324 (9217)

PT 95632 A 920430 (9222)#

JP 04506656 W 921119 (9301)

12 pp

ES 2038086 A6 930701 (9331)#

AU 639232 B 930722 (9336)

ADT ZA 9008391 A ZA 90-8391 901019; EP 469010 A EP 90-906144 900419; JP  
 04506656 W JP 90-506312 900419, WO 90-DK104 900419; ES 2038086 A6 ES  
 90-2611 901017; AU 639232 B AU 90-55278 900419

FDT JP 04506656 W Based on WO 9012561; AU 639232 B Previous Publ. AU  
 9055278, Based on WO 9012561

PRAI DK 89-1918 890420; ZA 90-8391 901019; ES 90-2611 901017

AN 90-348243 [46] WPIDS

AB WO 9012561 A UPAB: 960129

Admin. of a therapeutically or prophylactically effective amt. of a  
 sulphated mono, di- or oligosaccharide or their deriv, salt or  
 complex to a patient. Also claimed is the use of the sulphated  
 saccharide derivs for combating or preventing hair loss and/or  
 preserving natural colour of the hair.

Pref. embodiments - The mono saccharide is selected from  
 xylose, fructose and glucose. and the disaccharide is selected from  
 sucrose, lactose, maltose and cellobiose. The disaccharide deriv.  
 is selected from sucrose pentasulphate, sucrose hexasulphate,  
 sucrose heptasulphate and sucrose sulphate and is in the form of or  
 Na salt (esp surcalyate). The saccharide deriv. is combined with a  
 glucosamino-glycan selected from **hyaluronic** acid, dermatan  
 sulphate, chondroitin sulphate, keratan sulphate, heparan and  
 heparan sulphate.

USE/ADVANTAGE - The saccharide derivs have been indicated for  
 alleviating symptoms of anorectal disease and for promoting wound  
 healing. The saccharide derivs. are used in concs. of 0.5 - 15 wt.%.  
 The compsn. is **topically** applied in the form of ointment,  
 lotion, gel, cream, emulsa. soln. shampoo, soap, spray, paste,  
 powder, sponge, hair tonic etc. or injected or introduced

parenterally or **implanted** into the scalpetic. @ (30pp

Dwg.No.0/0)

0/0

ABEQ JP04506656 W UPAB: 930928

Admin. of a therapeutically or prophylactically effective amt. of a  
 sulphated mono, di- or oligosaccharide or their deriv., salt or  
 complex to a patient. Also claimed is the use of the sulphated  
 saccharide derivs for combating or preventing hair loss and/or  
 preserving natural colour of the hair.

The monosaccharide is pref. selected from JP4506656A - Wm xylose,  
 fructose and glucose, and the disaccharide is selected from sucrose,  
 lactose, maltose and cellobiose. The disaccharide deriv. is selected  
 from sucrose pentasulphate, sucrose hexasulphate, sucrose  
 heptasulphate and sucrose sulphate and is in the form of or Na salt  
 (esp surcalyate). The saccharide deriv. is combined with a

glucosamino-glycan selected from **hyaluronic** acid, dermatan sulphate, chondroitin sulphate, keratin sulphate, heparan and heparan sulphate.

USE/ADVANTAGE - The saccharide derivs. have been indicated for alleviating symptoms of anorectal disease and for promoting wound healing. The saccharide derivs. are used in concns. of 0.5-15 wt.%. The compsn. is **topically** applied in the form of ointment, lotion gel, cream, emulsa, soln. shampoo, soap, spray, paste, powder, sponge, hair tonic etc. or injected or introduced parenterally or **implanted** into the scalpetic

L86 ANSWER 25 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD  
 AN 89-278153 [38] WPIDS  
 DNC C89-123136  
 TI **Topical** treatment of teeth and supporting tissue - using sulphated saccharide esp. poly sulphated or per sulphated saccharide.  
 DC B03 D21  
 IN BAR-SHALOM, D; BUKH, N; HAMBURGER, J; BAR-SHALOM, D K; BARSHALOM, D  
 PA (BUKH-N) BUKH MEDITEC AS; (TAND-N) TANDLAEGESELKABET HAMBURGER APS  
 JESPER; (BUKH-N) BUKH AS NIELS; (BUKH-I) BUKH N; (BUKH-N) BUKH N  
 A/S; (NIEL-N) NIELS BUKH A/S; (TAND-N) TANDLAEGESELKABET HAMBU  
 CYC 36  
 PI WO 8907932 A 890908 (8938)\* EN 44 pp  
 RW: AT BE CH DE FR GB IT LU NL OA SE  
 W: AT AU BB BR CH DE DK FI GB HU JP KP KR LK LU MC MG MW NL NO  
 RO SD SE SU US  
 AU 8940744 A 890922 (8950)  
 ZA 8906525 A 900530 (9026)#  
 EP 404792 A 910102 (9102)  
 R: AT BE CH DE FR GB IT LI LU NL SE  
 DK 9002043 A 900824 (9107)  
 PT 91547 A 910418 (9118)#  
 ES 2018728 A 910501 (9123)#  
 CN 1049608 A 910306 (9145)#  
 JP 05503280 W 930603 (9327) 15 pp  
 US 5240710 A 930831 (9336) 13 pp  
 EP 404792 B1 931020 (9342) EN 21 pp  
 R: AT BE CH DE FR GB IT LI LU NL SE  
 DE 68910116 E 931125 (9348)  
 DK 169606 B 941227 (9506)  
 IL 91438 A 950330 (9530)#  
 CA 1336682 C 950815 (9542)#  
 ADT WO 8907932 A WO 89-DK43 890224; ZA 8906525 A ZA 89-6525 890825; EP  
 404792 A EP 89-903119 890224; ES 2018728 A ES 89-2938 890825; JP  
 05503280 W JP 89-502933 890224; WO 89-DK43 890224; US 5240710 A Cont  
 of US 89-375006 890804; US 92-939969 920904; EP 404792 B1 EP  
 89-903119 890224; WO 89-DK43 890224; DE 68910116 E DE 89-610116  
 890224; EP 89-903119 890224; WO 89-DK43 890224; DK 169606 B WO  
 89-DK43 890224; DK 90-2043 900824; IL 91438 A IL 89-91438 890825; CA  
 1336682 C CA 89-609143 890823  
 FDT JP 05503280 W Based on WO 8907932; EP 404792 B1 Based on WO 8907932;  
 DE 68910116 E Based on EP 404792, Based on WO 8907932; DK 169606 B  
 Previous Publ. DK 9002043  
 PRAI DK 88-5055 880909; DK 88-1024 880226; ZA 89-6525 890825;  
 DK 90-2043 900824; ES 89-2938 890825; IL 89-91438 890825;  
 CA 89-609143 890823  
 AN 89-278153 [38] WPIDS  
 AB WO 8907932 A UPAB: 951004

Use of a sulphated saccharide (I) or its salt or complex as an ingredient in a **topical** prepn. for the prophylaxis or treatment of diseases or conditions of the teeth or tooth-supporting tissue in partic. plaque-related conditions is new.

The saccharide is pref. monosaccharide such as xylose, fructose, or glucose, a dissacharide such as sucrose, lactose, maltose, or cellulose, or a polysaccharide such as dextran, heparon, dermatan, proteodermaton, **hyaluronic** acid, heporin, chondroitin, amylose, glucosamine, glucosaminoglycon or a mucopolysaccharide or subunit. (I) may be complexed with or forms a

salt with an alkaline earth metal (such as Na, K, Ca, Sr, Mg, or Ba), Al, Ga, Zn, Cu, B, Mn or an organic base (e.g. an amino acid) esp. Al opt. as aluminium hydroxide. (I) is esp. a sodium and/or potassium salt of sucrose octakis (hydrogen sulphate) or is is sucralfate.

USE - Diseases or conditions which can be treated include dental canes, dental plaque, gingivitis, periodontitis, alveolitis, pulpitis, osteomyelitis, post-extractive or post-surgical wounds, tooth eruption, bone resorption, prosthetic irritation, cysts, or neoplasms originating in the tooth, supporting tissue, and bacterial mycotic and viral oral **infections**.

0/0

Dwg.0/0

ABEQ JP05503280 W UPAB: 931116

Use of a sulphated saccharide (I) or its salt or complex as an ingredient in a **topical** prepn. for the prophylaxis or treatment of diseases or conditions of the teeth or tooth-supporting tissue in partic. plaque-related conditions is new.

The saccharide is pref. monosaccharide such as xylose, fructose, or glucose, a disaccharide such as sucrose, lactose, maltose or cellulose, or a polysaccharide such as dextran, heparon, dermatan, proteodermaton, **hyaluronic** acid, heparin, chondroitin, amylase, glucosamine, glucosaminoglycon or a mucopolysaccharide or subunit. (I) may be complexed with or forms a salt with an alkaline earth metal (such as Na, K, Ca, Sr, Mg, or Ba), Al, Ga, Zn, Cu, B, Mn or an organic base (e.g. an amino acid) esp. Al opt. as aluminium hydroxide. (I) is esp. a sodium and/or potassium salt of sucrose octakis (hydrogen sulphate) or is sucralfate.

USE - Diseases or conditions which can be treated include dental caries, dental plaque, gingivitis, periodontitis, alveolitis, pulpitis, osteomyelitis, post-extractive or post-surgical wounds, tooth eruption, bone resorption, prosthetic irritation, cysts, or neoplasms originating in the tooth, supporting tissue, and bacterial mycotic and viral oral **infections**.

ABEQ US 5240710 A UPAB: 931122

Dental disease in a human is prevented, diminished or treated, by administering a salve, paste, gel or cream prepn. contg. a prophylactic or therapeutic amt. of Al-salt or complex of sulphated saccharide.

Pref. saccharide is poly- or persulphated sucrose, lactose, maltose or cellulose. Al-cpd. is Al(OH)<sub>3</sub> or aluminium disaccharide polysulphate.

ADVANTAGE - Compsn. can also comprise an adhesive which adheres to the teeth or tooth-supporting tissue.

Dwg.0/0

ABEQ EP 404792 B UPAB: 931202

Use of an aluminium complex of sulphated sucrose as an ingredient for the manufacture of a **topical** preparation for the prophylaxis or treatment of diseases or conditions of the teeth or tooth-supporting tissue selected from dental caries, dental plaque, gingivitis, periodontitis, alveolitis, pulpitis, post-extractive or post-surgical wounds, tooth eruption, bone resorption, prosthetic irritation, cysts or neoplasms originating in the tooth-supporting tissue, and bacterial, mycotic and viral oral **infections**, in particular for the prophylaxis or treatment of inflammatory and plaque-related conditions.

Dwg.0/0

L86 ANSWER 26 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD

AN 89-206453 [28] WPIDS

CR 89-206452 [28]

DNC C89-091672

TI **Topical** compsn. comprising sulphated saccharide - for application to skin or non-gastrointestinal, non-oral, non-bladder mucosa to treat e.g. inflammation, burns, irritation, etc..

DC B03 B04 C02 C03

IN BAR-SHALOM, D; BUKH, N; BARSHALOM, D; BURKH, N

PA (BARS-I) BAR-SHALOM D; (BUKH-N) BUKH MEDITEC AS; (BUKH-N) BUKH

MEDITEC A/S; (BUKH-N) BUKH MEDITEK AS

CYC 32

PI WO 8905646 A 890629 (8928)\* EN 43 pp  
 RW: AT BE CH DE FR GB IT LU NL OA SE  
 W: AT AU BB BG BR CH DE DK FI GB HU JP KP KR LK LU MC MG MW NL  
 NO RO SD SE SU US  
 AU 8929146 A 890719 (8941)  
 DK 9001515 A 900814 (9044)  
 EP 394333 A 901031 (9044)  
 R: AT BE CH DE FR GB IT LI LU NL SE  
 CA 2020199 A 911230 (9213)#  
 JP 04500798 W 920213 (9213) 16 pp  
 DK 9200057 A 920117 (9229)  
 AU 9333960 A 930506 (9325)  
 KR 9303117 B1 930419 (9420)  
 EP 394333 B1 950315 (9515) EN 10 pp  
 R: AT BE CH DE FR GB IT LI LU NL SE  
 DE 3853365 G 950420 (9521)  
 JP 07039347 B2 950501 (9522) 12 pp  
 AU 664419 B 951116 (9602)

ADT WO 8905646 A WO 88-DK217 881221; AU 8929146 A AU 89-29146 881221; DK 9001515 A DK 90-1515 900621; EP 394333 A EP 89-901102 881221; CA 2020199 A CA 90-2020199 900629; JP 04500798 W JP 89-501022 881221; DK 9200057 A Div ex DK 90-1515 881221, DK 92-57 920117; AU 9333960 A Div ex AU 89-29146 881221, AU 93-33960 930303; KR 9303117 B1 WO 88-DK217 881221, KR 89-701562 890821; EP 394333 B1 WO 88-DK217 881221, EP 89-901102 881221; DE 3853365 G DE 88-3853365 881221, WO 88-DK217 881221, EP 89-901102 881221; JP 07039347 B2 WO 88-DK217 881221, JP 89-501022 881221; AU 664419 B Div ex AU 89-29146 881221, AU 93-33960 930303

FDT EP 394333 B1 Based on WO 8905646; DE 3853365 G Based on EP 394333, Based on WO 8905646; JP 07039347 B2 Based on JP 04500798, Based on WO 8905646; AU 664419 B Previous Publ. AU 9333960

PRAI DK 87-6740 871221; DK 88-5054 880909; WO 88-DK217 881221

AN 89-206453 [28] WPIDS

CR 89-206452 [28]

AB WO 8905646 A UPAB: 950404  
 Compsn., partic. for **topical** applicn. to skin or any non-gastrointestinal, non-oral, non-bladder mucosal surface comprises a sulphated saccharide (I) or salt or complex, with an acceptable carrier or excipient. A non-sulphated polysaccharide eg **hyaluronic acid**, may also be present.  
 USE - Used for preventing or treating non-bladder premalignant or malignant disorders; for preventing or treating irritation or burns of the skin, connective tissue or non-oral mucosa; for preventing or treating skin, connective tissue or mucosal ageing; or for preventing or treating **infectious**, malignant or allergic/ immune disorders (all claimed).  
 (I) may be used in tissue culture media (claimed) and for coating eg. catheters to reduce thrombus formation or prevent inflammatory responses.  
 Dwg.0/0  
 Dwg.0/0

ABEQ EP 394333 B UPAB: 950425  
 Use of sulphated mono- or disaccharide or a salt or complex thereof for combatting or preventing ageing of skin, including treating or preventing skin wrinkles.  
 Dwg.0/0

L86 ANSWER 27 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD

AN 89-206452 [28] WPIDS

CR 89-206453 [28]

DNC C89-091671

TI **Topical** pharmaceutical compsns. - contg. sucralfate, for treating skin and mucosal disorders.

DC B03 B07

IN BAR-SHALOM, D; BUKH, N; BARSHALOM, D

PA (BARS-I) BAR-SHALOM D; (BUKH-N) BUKH MEDITEC AS; (BUKH-N) BUKH MEDITEC A/S; (BUKH-N) BUKH MEDITEK AS



CYC 32  
 PI WO 8905645 A 890629 (8928)\* EN 40 pp  
 RW: AT BE CH DE FR GB IT LI LU NL OA SE  
 W: AT AU BB BG BR CH DE DK FI GB HU JP KP KR LK LU MC MG MW NL  
 NO RO SD SE SU US  
 AU 8929145 A 890719 (8941)  
 DK 9001516 A 900815 (9044)  
 EP 420849 A 910410 (9115)  
 R: AT BE CH DE FR GB IT LI LU NL SE  
 CA 2020200 A 911230 (9213)#  
 JP 04500797 W 920213 (9213) 13 pp  
 DK 9400203 A 940218 (9425)  
 DK 169018 B 940801 (9429)  
 EP 640346 A1 950301 (9513) EN  
 R: AT BE CH DE FR GB IT LI LU NL SE  
 ADT WO 8905645 A WO 88-DK216 881221; AU 8929145 A AU 89-29145 881221; DK  
 9001516 A DK 90-1516 900621; EP 420849 A EP 89-901101 881221; JP  
 04500797 W JP 89-501021 881221; DK 9400203 A Div ex DK 92-57 920117,  
 DK 94-203 940218; DK 169018 B Div ex DK 90-1515 900621, DK 92-57  
 920117; EP 640346 A1 Related to EP 89-901102 881221, EP 94-202490  
 881221  
 FDT DK 169018 B Previous Publ. DK 9200057  
 PRAI DK 87-6740 871221; DK 88-5054 880909; WO 88-DK217 881221;  
 DK 90-1516 900621; WO 88-DK217 881221  
 AN 89-206452 [28] WPIDS  
 CR 89-206453 [28]  
 AB WO 8905645 A UPAB: 950404  
 Pharmaceutical compsns, esp for **topical** application to  
 skin or non-bladder, non-gastrointestinal, non-oral mucosa, comprise  
 sucralfate (I) and a carrier or excipient. (I) is a sucrose  
 octasulphate Al complex (see US3432489).  
 The compsns pref contain 0.001-99 (esp 1-10) wt% (I), opt  
 together with a non-sulphonated polysaccharide, e.g.  
**hyaluronic acid**. (I) has a particle size of up to 200 (e.g.  
 1-5) microns. The compsns are formulated as powders, pastes,  
 ointments, lotions, gels, creams, salves, emulsions, suspensions,  
 sprays, sponges, strips, plasters, pads, dressings or ostomy plates,  
 and are applied 1-10 times a day.  
 USE - The compsns may be applied to the skin, lips, perianal  
 areas, nose, respiratory tract, eyes, ears, vagina or vulva for  
 treatment or prophylaxis of inflammations, **infections**,  
 irritations, burns, ulcers, wounds and pre-malignant or malignant  
 disorders, for modification of tissue regeneration, for modulation  
 of immune reactions and for combatting ageing.  
 Dwg.0/0  
 Dwg.0/0

L86 ANSWER 28 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD  
 AN 89-009088 [02] WPIDS  
 DNC C89-004216  
 TI Compsn. to prevent fibrin deposition or adhesion formation -  
 comprising sparingly soluble enzyme, esp. tissue plasminogen  
 activator, for **topical** application.  
 DC B04 C03  
 IN MOHLER, M A; NGUYEN, T H  
 PA (GETH) GENENTECH INC  
 CYC 25  
 PI EP 297860 A 890104 (8902)\* EN 20 pp  
 R: AT BE CH DE ES FR GB GR IT LI LU NL SE  
 WO 8900049 A 890112 (8905) EN  
 W: AU DK FI HU JP KR NO  
 AU 8819985 A 890130 (8920)  
 PT 87886 A 890630 (8930)  
 DD 271268 A 890830 (9006)  
 FI 8906359 A 891229 (9012)  
 ZA 8804739 A 900328 (9017)  
 NO 8905338 A 900402 (9019)  
 DK 8906753 A 900228 (9020)  
 HU 56283 T 910828 (9138)

3-18

JP 04502753 W 920521 (9227) 19 pp  
 EP 297860 B1 930901 (9335) EN 24 pp  
 R: CH DE FR GB LI  
 IL 86933 A 930708 (9335)  
 DE 3883645 G 931007 (9341)  
 IL 102625 A 930922 (9349)  
 HU 209955 B 941228 (9506)  
 IE 65658 B 951115 (9605)  
 ADT EP 297860 A EP 88-305935 880630; WO 8900049 A WO 88-US2194 880630;  
 ZA 8804739 A ZA 88-4739 880701; JP 04502753 W JP 88-505978 880630,  
 WO 88-US2194 880630; EP 297860 B1 EP 88-305935 880630; IL 86933 A IL  
 88-86933 880630; DE 3883645 G DE 88-3883645 880630, EP 88-305935  
 880630; IL 102625 A IL 88-102625 880630; HU 209955 B HU 88-4201  
 880630, WO 88-US2194 880630; IE 65658 B IE 88-2023 880701  
 FDT JP 04502753 W Based on WO 8900049; DE 3883645 G Based on EP 297860;  
 IL 102625 A Div ex IL 86933; HU 209955 B Previous Publ. HU 56283,  
 Based on WO 8900049  
 PRAI US 87-68872 870701; US 87-125319 871125; US 88-210895 880624  
 AN 89-009088 [02] WPIDS  
 AB EP 297860 A UPAB: 930923

A pharmaceutical compsn. to prevent fibrin deposition or adhesion formation that is **topically** applicable and capable of delivering an enzyme for from 3 days to 2 weeks comprises a therapeutically effective amt. of a sparingly soluble enzyme. More specifically the enzyme is tissue plasminogen activator (+PA). The compsn. may also include an inert, adherence enhancing vehicle, e.g. petrolatum jelly or **hyaluronic** acid.

Also claimed is a dispensing device for the admin. of a pharmaceutical compsn. to prevent fibrin deposition or adhesion formation comprising (a) a first container contg. a sparingly soluble enzyme and (b) a second container contg. an adherence enhancing vehicle, at least one of the containers being flexible. Also claimed is the use of a sparingly soluble enzyme in the mfr. of a compsn. for **topical** application for preventing, ameliorating or reversing fibrin deposition.

USE/ADVANTAGE - The compsns. are used for preventing formation or reformation of adhesions, partic. in the peritoneal or pelvic cavities resulting from surgery, **infection**, inflammation or trauma. The enzyme dissolves slowly over a period of time enabling a single **topical** application to provide continuous release of active enzyme.

0/3

ABEQ EP 297860 B UPAB: 931119  
 A **topically** applicable pharmaceutical compsn. to prevent fibrin deposition or adhesion formation, comprising a therapeutically effective amount of tissue plasminogen activator ('t-PA') in a sparingly soluble solid form which dissolves at a desired rate in a biofluid such that said composition is capable of delivering t-PA for a period from 3 days to two weeks when applied to a site of potential fibrin deposition or adhesion formation consequent on surgery, the rate of dissolution in use being due to the sparing solubility of the t-PA.  
 Dwg.0/3

L86 ANSWER 29 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD  
 AN 87-362420 [51] WPIDS  
 DNC C87-155254  
 TI Vitamin-C, zinc salt and sulphur contg. amino-acid compsn. - for **topical** treatment of skin conditions by stimulating epithelial tissue.

DC B05  
 IN FAHIM, M S  
 PA (FAHI-I) FAHIM M S  
 CYC 12  
 PI US 4711780 A 871208 (8751)\* 17 pp  
 EP 314835 A 890510 (8919) EN  
 R: BE CH DE FR GB IT LI LU NL SE  
 CA 1291034 C 911022 (9149)#  
 EP 314835 B 920429 (9218) EN 17 pp

R: BE CH DE FR GB IT LI LU NL SE

DE 3778703 G 920604 (9224)#

ADT US 4711780 A US 86-862051 860512; EP 314835 A EP 87-116429 871106;  
 EP 314835 B EP 87-116429 871106; DE 3778703 G DE 87-3778703 871106  
 PRAI US 86-862051 860512; EP 87-116429 871106  
 AN 87-362420 [51] WPIDS  
 AB US 4711780 A UPAB: 930922

Medicament comprises vitamin C, a zinc salt, a sulphur contg. amino acid and opt. a polysaccharide or mucopolysaccharide.

Specifically the sulphur contg. amino acid is cysteine, cystine or methionine, the zinc salt is the sulphate and the polysaccharide is chondroitin sulphate, **hyaluronic acid**, calcium heparinate, dermatan sulphate or keratin sulphate.

USE - The medicament has a wide variety of applications such as the treatment of vaginitis, cervicitis, urethral **infections**, irritated bladder, extropian eyelids, blepharitis, keratitis, pink eye, burns, cuts, fever blisters, poison ivy wheals, insect bites, nappy rash, genital herpes, sunburn.

0/2

ABEQ DE 3778703 G UPAB: 930922

Medicament comprises vitamin C, a zinc salt, a sulphur contg. amino acid and opt. a polysaccharide or mucopolysaccharide.

Specifically the sulphur contg. amino acid is cysteine, cystine or methionine, the zinc salt is the sulphate and the polysaccharide is chondroitin sulphate, **hyaluronic acid**, calcium heparinate, dermatan sulphate or keratin sulphate.

USE - The medicament has a wide variety of applications such as the treatment of vaginitis, cervicitis, urethral **infections**, irritated bladder, extropian eyelids, blepharitis, keratitis, pink eye, burns, cuts, fever blisters, poison ivy wheals, insect bites, nappy rash, genital herpes, sunburn.

ABEQ EP 314835 B UPAB: 930922

A composition for treating epithelial tissue characterised by vitamin C in an amount from 0.5 to 30% by weight, a zinc salt present in an amount from 0.25 to 20% by weight, heptahydrate or the equivalent amount of zinc present as some other zinc salt, and a sulphur amino acid in an amount from 0.25 to 5% by weight.

L86 ANSWER 30 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD

AN 87-277600 [39] WPIDS

DNC C87-117936

TI New **hyaluronic acid** heavy metal salts - useful as antibacterials esp. the silver salt, for treating rheumatoid arthritis esp. as gold salt and in diagnosis when radio-labelled.

DC B04 D22 K08

IN GREENMAN, B; NIMROD, A

PA (BIOT-N) BIO-TECHN GEN CORP

CYC 17

PI WO 8705517 A 870924 (8739)\* EN 52 pp

RW: AT BE CH DE FR GB IT LU NL SE

W: AU DK JP

AU 8772068 A 871009 (8751)

EP 259485 A 880316 (8811) EN

R: AT BE CH DE FR GB IT LI LU NL SE

DK 8705974 A 871113 (8812)

US 4746504 A 880524 (8823) 11 pp

JP 63502670 W 881006 (8846)

US 4784991 A 881115 (8848) 12 pp

IL 81877 A 910310 (9120)

CA 1291123 C 911022 (9149)

ADT WO 8705517 A WO 87-US549 870313; EP 259485 A EP 87-902255 870313; US 4746504 A US 86-840419 860314; JP 63502670 W JP 87-502147 870313; US 4784991 A US 87-23666 870309

PRAI US 86-840419 860314; US 87-19474 870226; US 87-23666 870309

AN 87-277600 [39] WPIDS

AB WO 8705517 A UPAB: 930922

**Hyaluronic acid** heavy metal salts (I) are new, esp. the Ag, Au, Ce or W salts. The acid may be radioactively labelled, esp. with carbon-14.

USE/ADVANTAGE - (I) have various therapeutic uses. The Ag salt inhibits microbial growth and is suitable for **topical** admin. to burns, wounds, soft-tissue **infections**, ophthalmological **infections**, sepsis, keratitis etc. opt. in conjunction with an antibiotic. The Au salt on intra-articular admin. is useful for treating rheumatoid arthritis, joint inflammation etc. The radioactively labelled (I) can be used for diagnostic purposes. (I) may also be used in deodorants, cosmetic creams, lotions and sprays. The metal ions are slowly released from (I).

0/1

ABEQ US 4746504 A UPAB: 930922

Heavy metal salt (pref. Ag, Au, Ce, W); esp. Ag) of **hyaluronic** acid, is new. It may be prepd. by mixing aq. Na hyaluronate with molar excess AgNO<sub>3</sub>, then pptn. and recovery of Ag hyaluronate viz. by centrifugation, washing with ethanol, drying over N<sub>2</sub>, then vacuum drying, in dark.

USE - As antimicrobials, e.g., **topically** for burns and wounds and esp. to treat arthritis by intra-articular admin. and in C14-labelled form for diagnosis. **Hyaluronic** acid is 50-1500KD glucosamineglycan, natural or synthetic, as viscous soln., without immuno reactions and gives slow in vivo release of Ag,+.

ABEQ US 4784991 A UPAB: 930922

Heavy metal (i.e. Ag, Au, Ce, and W) salts or **hyaluronic** acid are new. Prepn. of these salts comprises addn. of a soluble metal salt to aq. Na hyaluronate; and pptn. with EtOH. Pref. prod. is silver hyaluronate.4 USE - The prods. are antimicrobial agents, and gold hyaluronate is a therapeutic for arthritis.

L86 ANSWER 31 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD

AN 87-088138 [13] WPIDS

CR 87-062629 [09]

TI New total and partial **hyaluronic** acid ester(s) - useful as medicaments, in cosmetics, as vehicles for medicines, in surgical articles etc..

DC A96 B04 B07 D21 D22 F01 F07 P34

IN DELLAVALLE, F; ROMEO, A; VALLE, F D; DELLA, VALLE F

PA (FIDI-N) FIDIA SPA

CYC 20

PI EP 216453 A 870401 (8713)\* EN 129 pp

R: AT BE CH DE FR GB IT LI LU NL SE

AU 8659836 A 870226 (8713)

NO 8602734 A 870202 (8713)

FI 8602878 A 870109 (8714)

JP 62064802 A 870323 (8717)

DK 8603236 A 870109 (8727)

HU 42512 T 870728 (8733)

ES 2001512 A 880601 (8922)

US 4851521 A 890725 (8937) 29 pp

FI 8902710 A 890602 (8945)

FI 8902711 A 890602 (8945)

FI 9001341 A 900316 (9022)

US 4965353 A 901023 (9045)

NO 9100295 A 870109 (9120)

IT 1203815 B 890223 (9125)

FI 9102618 A 910531 (9133)

FI 9102619 A 910531 (9133)

US 5202431 A 930413 (9317) 28 pp

US 5336767 A 940809 (9431) 28 pp

NO 175716 B 940815 (9432)

FI 94766 B 950714 (9534)

FI 94767 B 950714 (9534)

FI 94778 B 950714 (9534)

IL 79362 A 950731 (9540)

ADT EP 216453 A EP 86-305233 860707; AU 8659836 A AU 86-59836 860708; JP 62064802 A JP 86-161769 860708; ES 2001512 A ES 86-1101 860708; US 4851521 A US 86-881454 860702; US 4965353 A US 89-339919 890419; US 5202431 A Div ex US 86-881454 860702, Div ex US 89-339919 890419, Div ex US 90-562267 900803, Cont of US 91-663324 910301, US

91-794703 911120; US 5336767 A Div ex US 86-881454 860702, Div ex US 89-339919 890419, Div ex US 90-562267 900803, Div ex US 91-663324 910301, Div ex US 91-794703 911120, US 92-998749 921230; NO 175716 B NO 86-2734 860707; FI 94766 B FI 86-2878 860708; FI 94767 B Div ex FI 86-2878 860708, FI 90-1341 900316; FI 94778 B Div ex FI 86-2878 860708, FI 89-2710 890602; IL 79362 A IL 86-79362 860708

FDT US 5202431 A Div ex US 4851521, Div ex US 4965353; US 5336767 A Div ex US 4851521, Div ex US 4965353, Div ex US 5202431; NO 175716 B Previous Publ. NO 8602734; FI 94766 B Previous Publ. FI 8602878; FI 94767 B Previous Publ. FI 9001341; FI 94778 B Previous Publ. FI 8902710

PRAI IT 85-48322 850708; IT 86-48202 860630

AN 87-088138 [13] WPIDS

CR 87-062629 [09]

AB EP 216453 A UPAB: 940928

Total and partial esters (I) of **hyaluronic** acid with aliphatic araliphatic, cycloaliphatic or heterocyclic alcohols, and their salts, with (in)organic bases, except for the total Me ester of **hyaluronic** acid, are new. Pharmaceutical prepn. contg. as active ingredient (I) or its salt, and including the total Me ester of **hyaluronic** acid, is new.

USE/ADVANTAGE - (I) are useful in the formulation of, or as themselves, medicaments, in cosmetics (esp. with alcohols used in perfumery) and in medical veterinary, and surgical articles and prepn. (I) may be dominated by the properties of the **hyaluronic** acid or by the properties of the alcohol and/or salt component, e.g. when derived from steroid alcohols, the esters have activities such as anti-inflammatory activity with a better balanced, constant and regular action and prolonged release. Better stability and bioavailability, desired solvent solubility etc. may also be achieved. The (I) are typically, vehicles for anaesthetics, analgesics, anti-inflammatory agnets, vasoconstrictors, antibacterials and antivirals, esp. for **topical** use, e.g. in ophthalmology, dermatology, etc. (I) may be in film form, to replace skin, as capsuled or microcapsules for subcutaneous **implantation**, as solid insert, as sponge for application to wounds, as threads to be woven into gauzes or used as sutures.

Dwg.0/0

Dwg.0/0

ABEQ US 4851521 A UPAB: 930922

Total and partial **hyaluronic** acid esters with aliphatic, araliphatic, cycloaliphatic and heterocyclic alcohols and salts, are new. Alcohols have up to 34C atoms and are opt. substd. 1 or 2 functional gps. and with C atoms opt. interrupted with O, S or N atoms, and include steroid cortisones, streptomycin, etc..

New process for their prod., comprises treating quat. NH<sub>4</sub> salt of polysaccharide (e.g. obtd. from cock's combs with esterifying agent in aprotic solvent and appropriate salification. Various MW fractions up to 13 million.

USE - Biocompatible vehicle and active cpds. in medicine, surgery, cosmetics, artificial skin, suture threads, etc..

ABEQ US 4965353 A UPAB: 930922

**Hyaluronic** esters in which all or only some of the COOH gps. have been esterified and nontoxic salts of the partial esters are new. Typical esters are benzyl, n-propyl and ethyl esters. Prepn. of these esters comprises condensn. of corresp. tetraalkylammonium salts with benzyl or alkyl halides in dimethyl sulphoxide.

USE - The prods. are converted to threads, gauzes, sponges, films and microcapsules, for medicinal, pharmaceutical and cosmetic applications.

ABEQ US 5202431 A UPAB: 931025

Partial eser of **hyaluronic** acid with an alcohol of the aliphatic, araliphatic, cycloaliphatic or heterocyclic sereis, has an at least a first portion of carboxylic acid gps. of the **hyaluronic** acid are salified with therapeutically active amione.

Pref. amine is alkaloids, peptides, phenothaiziones, benzodiazepines, thiosanthenes, hormones, vitamins,

anti-convulsants, anti-psychotics, anti-emetics, anaesthetic, hypnotics, anorexics, antibacterials, antivirals, anti-malarials, narcotic antagonists, antiinflammatory agents. The partial ester is pref. in a microcapsule.

USE - Used in pharmaceutical and cosmetic fields and biodegradable plastic materials.

Dwg. 0/0

ABEQ US 5336767 A UPAB: 940921

**Hyaluronic** esters and/or partial esters in which the alcoholic component is cortisone, hydrocortisone, fluorocortisone, corticosterone, deoxycorticosterone, prednisolone, prednisone, dexamethasone, betamethasone, paramethasone, flumethasone, fluocinolone, or its acetonide, fluprednylidene, clobetasol or beclomethasone, and their nontoxic salts are new.

USE/ADVANTAGE - These ester derivs. have bioplastic and pharmaceutical properties for medical, surgical and cosmetic applications. The prods. are therapeutics, for damaged skin, tendons, tissues, muscles and cartilage, and support tissue hydration, lubrication, cell migration, cell functions and cell differentiation, etc.

Dwg. 0/0

L86 ANSWER 32 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD

AN 86-271924 [42] WPIDS

CR 83-14905K [07]; 85-099097 [17]; 86-162168 [25]; 88-014163 [02]

DNC C86-117920

TI **Topical** compsn. contg. **hyaluronic** acid deriv. as vehicle - esp. for ophthalmic and dermatological application and new barium hyaluronate salts.

DC A96 B04 C03

IN DELLA, VALLE F; LORENZI, S; ROMEO, A; VALLE, D; VALLE, F D

PA (FIDI-N) FIDIA SPA; (FIDI-N) FIDIA FARM ITAL DERIVATI IND

CYC 22

PI BE 904547 A 861003 (8642)\* 58 pp

EP 197718 A 861015 (8642) EN

R: AT DE GB IT NL SE

FR 2579895 A 861010 (8647)

AU 8655662 A 861016 (8648)

JP 61236732 A 861022 (8649)

NO 8601331 A 861027 (8650)

PT 82342 A 861105 (8650)

LU 86386 A 860902 (8651)

FI 8601395 A 861006 (8703)

HU 40579 T 870128 (8710)

DK 8601498 A 861006 (8726)

ZA 8602463 A 871005 (8804)

ES 8800055 A 880101 (8809)

US 4736024 A 880405 (8816)

CH 672886 A 900115 (9007)

IT 1184675 B 871028 (9041)

IT 1229075 B 910717 (9232)

US 5166331 A 921124 (9250) 26 pp

EP 555898 A2 930818 (9333) EN 30 pp

R: AT DE GB IT NL SE

EP 197718 B1 931215 (9350) EN 37 pp

R: AT DE GB IT NL SE

DE 3689384 G 940127 (9405)

IL 78263 A 931115 (9405)

NO 174277 B 940103 (9406)

HU 208833 B 940128 (9409)

EP 555898 A3 931020 (9510)

US 5442053 A 950815 (9538) 29 pp

IE 64440 B 950809 (9539)

ADT BE 904547 A BE 86-904547 860403; EP 197718 A EP 86-302291 860327; FR 2579895 A FR 86-4601 860401; JP 61236732 A JP 86-79060 860404; ZA 8602463 A ZA 86-2463 860403; ES 8800055 A ES 86-553714 860404; US 4736024 A US 86-847632 860403; IT 1229075 B IT 85-47924 850405; US 5166331 A CIP of US 84-669431 841108, CIP of US 85-719113 850402, Cont of US 85-756824 850719, US 89-452681 891219; EP 555898 A2

Related to EP 86-302291 860327, EP 93-200175 860327; EP 197718 B1 EP 86-302291 860327; DE 3689384 G DE 86-3689384 860327, EP 86-302291 860327; IL 78263 A IL 86-78263 860325; NO 174277 B NO 86-1331 860404; HU 208833 B Div ex HU 86-1402 860403, HU 89-1006 860403; EP 555898 A3 EP 93-200175 860327; US 5442053 A CIP of US 82-425462 820928, CIP of US 84-669431 841108, CIP of US 85-719113 850402, Cont of US 85-756824 850719, Cont of US 89-452681 891219, US 92-931949 920819; IE 64440 B IE 86-847 860327

FDT DE 3689384 G Based on EP 197718; NO 174277 B Previous Publ. NO 8601331; US 5442053 A CIP of US 4593091, Cont of US 5166331

PRAI IT 85-48980 851223; IT 85-47924 850405; IT 83-49143 831010; IT 84-48979 841009

AN 86-271924 [42] WPIDS

CR 83-14905K [07]; 85-099097 [17]; 86-162168 [25]; 88-014163 [02]

AB BE 904547 A UPAB: 951004

Topical compsn. comprises (1) at least one **topically** -active pharmaceutical (I) and (2) **hyaluronic acid** (II), or one of its mol. fractions, opt. in the form of a salt with alkali or alkaline earth metals, Al, NH<sub>4</sub> or with one or more active ingredients (esp. (I)). The pure Ba salts (A) of noninflammatory (II) fractions of mol.wts. 250000-350000; 50000-100000 or 500000-730000, free of (II) of mol.wt. below 30000, are new.

USE/ADVANTAGE - The compsns. are useful for treating ophthalmic and dermatological disorders, and diseases of the mucosa, oral and nasal cavities, and outer ear, particularly in paediatric and veterinary medicine. The use of (II) provides better biological better biological availability than known vehicles. (II) particularly well tolerated by the corneal epithelium without risk of sensitisation and with a long-lasting adherence.

Dwg.0/0

Dwg.0/0

ABEQ US 4736024 A UPAB: 930922

The prepn. of a salt of **hyaluronic acid** with pharmacologically active substance comprises (a) combining aq. soln. of Ba salt of **hyaluronic acid** with sulphate of drug and (b) sepn. of BaSO<sub>4</sub> to give a salt of **hyaluronic acid** with drug in aq. soln. SO<sub>4</sub> and **hyaluronic acid** are stoichiometric, giving neutral salt, or partial salification.

Ba salt of **hyaluronic acid** may be further combined with sulphate of alkali, alkaline earth metal, Al or NH<sub>4</sub> stoichiometrically. A wide range of drugs may be used including antibiotics, erythromycin, etc. Mol.wt. fraction of

**hyaluronic acid** may be used viz. between 90-80% and 0.23% of M.W. of integral uronic acid i.e.  $13 \times 10^6$ , pref. none below 30000, (250000-350000). Cetylpyridinium salt of **hyaluronic acid** may be treated with BaCl<sub>2</sub> and Bahyaluronate pptd. with ethanol.

USE - Prepn. of **topical** and ophthalmic compsns. with increased bioavailability of drug.

ABEQ US 5166331 A UPAB: 930922

Pharmaceutical compsn. comprises as active ingredient a neutral or partially neutralised salt of **hyaluronic acid** or its MW fraction (Fig.1) with a basic drug for **topical** admin. readily absorbed intradermally or via the nasal or rectal mucosa, together with **topical** excipient.

Pref. a partial salt is used with an optical drug partially salified with alkali (ne earth) metal or Al or NH<sub>4</sub> as neutral salt. Mol. wt. fractions are 30000-730000, free of mol. wt. below 30000; 50000-100000; and 500000-730000.

Drugs include antibiotics, anti-infectives, antivirals, anti-inflammatory (NSAID's), wound healers, cytostatics, cytotoxics, anaesthetics, cholinergic promoters and antagonists for dermatological, otolaryngological, obstetrical and neurological use. Prepn. is by homogenisation of heñcrests in acetone, agitation, centrifugation and vacuum drying.

USE - 50000-100000 MW fractions for wound healing and 500000-730000 for intraocular and intra-articular injections without causing inflammation. HA enhances drug action.

0/1

ABEQ EP 555898 A UPAB: 931119

**Topical** compsn. comprises (1) at least one **topically-active** pharmaceutical (I) and (2) **hyaluronic** acid (II), or one of its mol. fractions, opt. in the form of a salt with alkali or alkaline earth metals, Al, NH<sub>4</sub> or with one or more active ingredients (esp. (I)). The pure Ba salts (A) of noninflammatory (II) fractions of mol.wts. 250000-350000; 50000-100000 or 500000-730000, free of (II) of mol.wt. below 30000, are new.

USE/ADVANTAGE - The compsns. are useful for treating ophthalmic and dermatological disorders, and diseases of the mucosa, oral and nasal cavities, and outer ear, particularly in paediatric and veterinary medicine. The use of (II) provides better biological availability than known vehicles. (II) particularly well tolerated by the corneal epithelium without risk of sensitisation and with a long-lasting adherence.

Dwg.0/0

ABEQ EP 197718 B UPAB: 940203

A medicament which comprises: (a) a pharmaceutically active substance or a mixture of pharmacologically active substances suitable for **topical** administration; and (b)

**hyaluronic** acid or a pharmaceutically acceptable salt of said **hyaluronic** acid, optionally together with an additional excipient suitable for **topical** administration, with the proviso that said active substance is not an ophthalmic drug when the **hyaluronic** acid is a fraction having an average molecular weight of from 50,000 to 730,000 and being substantially free of **hyaluronic** acid having a molecular weight of less than 30,000.

Dwg.0/0

ABEQ US 5442053 A UPAB: 950927

Partial or stoichiometrically neutral salt of **hyaluronic** acid (HA) or its mol.wt. fraction with at least one pharmacologically active substance (PAS) of a basic nature supplied for **topical** admin. is claimed.

The active substance is for dermatological, ophthalmological, otorhinolaryngological, odontological, angiological, obstetrical or neurological use as an antibiotic, **antiinfective**, antiviral, antimicrobial, antiinflammatory, wound healing, cytostatic cytotoxic, anaesthetic, cholinergic promoter, cholinergic antagonist, adrenergic promoter or adrenergic antagonist agent, e.g. kanamycin, amikacin, tobramycin, spectinomycin, oleandomycin, carbomycin, spiramycin, oxytetracycline, routetracycline, bacitracin, polymyxin B, gramicidin, colistin, chloramphenicol, uncomycin, amphotericin B, griseofulvin, myotatin, diethylcarbamazine, mebendazol, sulphacetamide, sulphadiazine, sulphisoxazole, iodoxuridine, adenine, arabinoside, trifluorothymidine, etc. Pref. the HA is a fraction of mol.wt. 30000-730000 (50000-100000) or 500000-730000.

USE/ADVANTAGE - The HA fractions are used e.g. for stimulating wound healing, for intraocular or intraarticular **infections** for replacing the endobulbar liqs. in the eye and for treating damaged bone joints, resp. The HA is pref. the vehicle in phthalmic solns. HA enhances the biological activity of ophthalmic drugs. The HA contg. compsns. have good tolerability to the cornea and allows the use of a high percentage of HA that can be obtd. from source tissues.

Dwg.1/1

L86 ANSWER 33 OF 33 WPIDS COPYRIGHT 1996 DERWENT INFORMATION LTD  
 AN 85-099097 [17] WPIDS  
 CR 83-14905K [07]; 86-162168 [25]; 86-271924 [42]; 88-014163 [02]  
 DNC C85-042889  
 TI New **hyaluronic** acid fractions - useful for wound healing or treating eye or joint disorders.  
 DC A96 B04 C03  
 IN DELLA, VALLE F; LORENZI, S; ROMEO, A  
 PA (DVAL-I) DELLA VALLE F; (FIDI-N) FIDIA SPA  
 CYC 24  
 PI BE 900810 A 850411 (8517)\* 37 pp



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FR 2553099 A 850412 (8520)

AU 8434148 A 850418 (8523)

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PT 79339 A 850509 (8526)

ZA 8407942 A 850404 (8528)

FI 8403990 A 850412 (8530)

DK 8404853 A 850412 (8531)

LU 85582 A 850604 (8541)

HU 36834 T 851028 (8601)

ES 8507573 A 851216 (8611)

JP 61028503 A 860208 (8612)

CA 1205031 A 860527 (8626)

KR 8601148 B 860818 (8652)

CN 85102921 A 861008 (8730)

CH 666897 A 880831 (8838)

EP 138572 B 900725 (9030)

R: AT DE GB IT NL SE

IT 1178041 B 870903 (9035)

DE 3482812 G 900830 (9036)

IL 73217 A 910730 (9133)

IT 1212892 B 891130 (9150)

IL 96943 A 930315 (9322)

JP 06008323 B2 940202 (9408)

US 5442053 A 950815 (9538) *dup* 29 pp

ADT BE 900810 A BE 84-900810 841011; EP 138572 A EP 84-306914 841010; FR 2553099 A FR 84-15547 841010; ZA 8407942 A ZA 84-7942 841011; JP 61028503 A JP 84-214046 841011; IL 96943 A IL 84-96943 841010; JP 06008323 B2 JP 84-214046 841011; US 5442053 A CIP of US 82-425462 820928, CIP of US 84-669431 841108, CIP of US 85-719113 850402, Cont of US 85-756824 850719, Cont of US 89-452681 891219, US 92-931949 920819

FDT IL 96943 A Div ex IL 73217; JP 06008323 B2 Based on JP 61028503; US 5442053 A CIP of US 4593091, Cont of US 5166331

PRAI IT 84-48979 841009; IT 83-49143 831011; IT 85-47924 850402

AN 85-099097 [17] WPIDS

CR 83-14905K [07]; 86-162168 [25]; 86-271924 [42]; 88-014163 [02]

AB BE 900810 A UPAB: 951004

Pure non-inflammatory **hyaluronic** acid fractions (I) with an av. molecular wt. of 30,000-730,000 and their Na and K salts are new. The fractions have molecular wts. of 50,000-100,000 (Ia), 250,000-350,000 (Ib) and 500,000-730,000 (Ic).

USE - (Ia) is useful for promoting wound healing. (Ic) is useful for treating disorders of the joints in humans and animals (esp. horses) and for replacing intra-ocular fluids. (Ib) is a combination of (Ia) and (Ib) and may be used for the same purpose as (Ia). (I) are also useful as carriers for drugs, e.g. pilocarpine, triamcinolone, epidermal growth factor, streptomycin or gentamicin. (Fl)

Dwg.0/1

Dwg.0/1

ABEQ EP 138572 B UPAB: 930925

A process for preparing a substantially pure, non-inflammatory, **hyaluronic** acid fraction comprising: subjecting starting material tissue to solvent extraction to produce a mixture containing **hyaluronic** acid, and subjecting the resulting mixture to molecular filtration to obtain a **hyaluronic** acid fraction having an average molecular weight of from 50,000 to 730,000, said fraction being substantially free of **hyaluronic** acid having a molecular weight of less than 30,000.

ABEQ US 5166331 A UPAB: 930925

Pharmaceutical compsn. comprises as active ingredient a neutral or partially neutralised salt of **hyaluronic** acid or its MW fraction (Fig.1) with a basic drug for **topical** admin. readily absorbed intradermally or via the nasal or rectal mucosa, together with **topical** excipient.

Pref. a partial salt is used with an optical drug partially

salified with alkali(ne earth) metal or Al or NH<sub>4</sub> as neutral salt. Mol. wt. fractions are 30000-730000, free of mol. wt. below 30000; 50000-100000; and 500000-730000.

Drugs include antibiotics, **anti-infectives**, antivirals, anti- inflammatories (NSAID's), wound healers, cytostatics, cytotoxics, anaesthetics, cholinergic promoters and antagonists for dermatological, otolinoloryngological, obstetrical and neurological use. Prepn. is by homogenisation of hencrests in acetone, agitation, centrifugation and vacuum drying.

USE - 50000-100000 MW fractions for wound healing and 500000-730000 for intraocular and intra-articular injections without causing inflammation. HA enhances drug action.

0/1

ABEQ US 5442053 A UPAB: 950927

Partial or stoichiometrically neutral salt of **hyaluronic** acid (HA) or its mol.wt. fraction with at least one pharmacologically active substance (PAS) of a basic nature supplied for **topical** admin. is claimed.

The active substance is for dermatological, ophthalmological, otorhinolaryngological, odontological, angiological, obstetrical or neurological use as an antibiotic, **antiinfective**, antiviral, antimicrobial, antiinflammatory, wound healing, cytostatic cytotoxic, anaepthetic, cholinergic promoter, cholinergic antagonist, adrenergic promoter or adrenergic antagonist agent, e.g. kanamycin, amikacin, tobramycin, spectinomycin, oleandomycin, carbomycin, spiramycin, oxytetracycline, routetracycline, bacitracin, polymyxin B, gramicidin, colistin, chloramphenicol, uncomycin, amphotericin B, griseofulvin, myotatin, diethylcarbamazine, mebendazol, sulphacetamide, sulphadiazine, sulphisoxazole, iodeoxuridine, adenine, arabinoside, trifluorothymidine, etc. Pref. the HA is a fraction of mol.wt. 30000-730000 (50000-100000) or 500000-730000.

USE/ADVANTAGE - The HA fractions are used e.g. for stimulating wound healing, for intraocular or intraarticular **infections** for replacing the endobulbar liqs. in the eye and for treating damaged bone joints, resp. The HA is pref. the vehicle in phthalmic solns. HA enhances the biological activity of ophthalmic drugs. The HA contg. compsns. have good tolerability to the cornea and allows the use of a high percentage of HA that can be obtd. from source tissues.

Dwg.1/1